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NEWS 3 Jul 12 BEILSTEIN enhanced with new display and select
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NEWS 4 resulting in a closer connection to BABS
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display fields
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and Japan
NEWS 6 AUG 02 Patent Office Classifications
The Analysis Edition of STN Express with Discover!
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NEWS 9 SEP 01 INPADOC: New family current-awareness alert (SDI)
available
NEWS 10 SEP 01 New pricing for the Save Answers for SciFinder
Wizard within
NEWS 11 SEP 01 STN Express with Discover!
WPIDS/WPINDEX/WPIX New display format, HITSTR, available in
NEWS 12 SEP 27 STANDARDS will no longer be available on STN
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***** STN Columbus *****

FILE 'HOME' ENTERED AT 14:24:22 ON 26 OCT 2004

=> index biosci
FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED
COST IN U.S. DOLLARS SINCE FILE
TOTAL ENTRY
SESSION
FULL ESTIMATED COST 0.42
INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOGIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 14:25:23 ON 26 OCT 2004

75 FILES IN THE FILE LIST IN STINDEX

Enter SET DETAIL ON to see search term postings or to view search error messages that display as 0* with SET DETAIL OFF.

=> fil reg
COST IN U.S. DOLLARS SINCE FILE
TOTAL ENTRY
SESSION
FULL ESTIMATED COST 1.14
1.56

FILE 'REGISTRY' ENTERED AT 14:26:32 ON 26 OCT 2004
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STRUCTURE FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6
DICTIONARY FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

SESSION
FULL ESTIMATED COST 0.57
3.05

FILE 'CAPLUS' ENTERED AT 14:28:02 ON 26 OCT 2004
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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/BMF
15 L1
S2155 BMF/RL
L3 0 L1/BMF
(L1 (L) BMF/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS SINCE FILE
TOTAL ENTRY
SESSION
FULL ESTIMATED COST 2.26
5.31

FILE 'CAPLUS' ENTERED AT 14:28:18 ON 26 OCT 2004
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Experimental and calculated property data are now available. For more information enter HELP PROP at an arrow prompt in the file or refer to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> S 300832-84-2/RN
L1 1 300832-84-2/RN

=> DEL SEL Y

=> SEL RN
E1 THROUGH E1 ASSIGNED

=> INDEX BEILSTEIN, GMELIN
COST IN U.S. DOLLARS SINCE FILE
TOTAL ENTRY

SESSION
FULL ESTIMATED COST 0.92
2.48

INDEX 'BEILSTEIN, GMELIN' ENTERED AT 14:27:43 ON 26 OCT 2004

2 FILES IN THE FILE LIST IN STINDEX

Enter SET DETAIL ON to see search term postings or to view search error messages that display as 0* with SET DETAIL OFF.

=> S E1 AND OPTICAL?/FA
FILE 'BEILSTEIN'
0 300832-84-2/BI
(300832-84-2/RN)
753186 OPTICAL?/FA
0 300832-84-2/BI AND OPTICAL?/FA
FILE 'GMELIN'
0 300832-84-2/BI
(300832-84-2/RN)
12899 OPTICAL?/FA
0 300832-84-2/BI AND OPTICAL?/FA

L2 QUE 300832-84-2/BI AND OPTICAL?/FA

=> DIS RANK
NO F-NUMBERS HAD GREATER THAN ZERO HITS

=> FIL CAPLUS
COST IN U.S. DOLLARS SINCE FILE
TOTAL ENTRY

strictly prohibited.

FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/BPN
15 L1
102210 BPN/RL
L4 0 L1/BPN
(L1 (L) BPN/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
7.57
SINCE FILE
ENTRY
2.26

FILE 'CAPLUS' ENTERED AT 14:28:28 ON 26 OCT 2004
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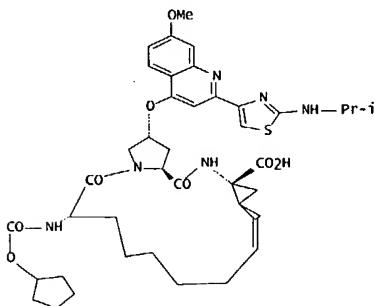
FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/IMF
15 L1
390491 IMF/RL
L5 0 L1/IMF
(L1 (L) IMF/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
7.57
SINCE FILE
ENTRY
2.26

KZ, MD; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG, TR.
(English). CODEN: PIXXD2. APPLICATION: WO 2003-US30402
20030925
PRIORITY: US 2002-PV414940 20020930; US 2002-PV421904 20021029;
US 2002-PV433834 20021216; US 2003-PV443662 20030130.
GI



AB Disclosed are oral pharmaceutical compns., kits and methods of treating and preventing Hepatitis C Viral (HCV) infections wherein Compound (I), a potent inhibitor of HCV serine protease, or a pharmaceutically acceptable salt thereof, is administered in a selected dosage range. Also disclosed are the use of I or a pharmaceutically acceptable salt thereof, as a control substance for validating an HCV replication assay and also as a control substance for determining the relative effectiveness of one or more substances, alone or in combination, to inhibit the replication of HCV.
IT 300832-84-2
or RL: PAC (Pharmacological activity); PEP (Physical, engineering chemical process); PYP (Physical process); THU (Therapeutic use);
BIOL (Biological study); PROC (Process); USES (Uses)
(potent inhibitor of HCV serine protease)

SESSION
FULL ESTIMATED COST
9.83
2.26

FILE 'CAPLUS' ENTERED AT 14:28:41 ON 26 OCT 2004
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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/PEP
15 L1
1687596 PEP/RL
L6 1 L1/PEP
(L1 (L) PEP/RL)

=> DIS L6 1 CBIB ABS HTRN

I6 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2004 ACS on STN
2004:310970 Document No. 140:327091 Potent inhibitor of HCV serine protease.

Chen, Shirlynn; Nehmiz, Gerhard; Croenlein, Jens Oliver; Steinmann, Gerhard; Gunn, Jocelyn Abella; Costa, Phuong Do (Boehringer Ingelheim International G.m.b.H., Germany). PCT Int. Appl. WO 2004030670

AI: 20040415, 42 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG,

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
14.95
SINCE FILE
ENTRY
5.12
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL
SESSION
CA SUBSCRIBER PRICE
0.70
SINCE FILE
ENTRY
-0.70

FILE 'CAPLUS' ENTERED AT 14:29:30 ON 26 OCT 2004
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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/PUR
15 L1
201134 PUR/RL
L7 0 L1/PUR
(L1 (L) PUR/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
17.21
SINCE FILE
ENTRY
2.26
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL
SESSION
ENTRY

CA SUBSCRIBER PRICE
0.70

0.00

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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S (L1/SPN OR L1/CPN)
15 L1
1660968 SPN/RL
6 L1/SPN
(L1 (L) SPN/RL)
15 L1
1155 CPN/RL
0 L1/CPN
(L1 (L) CPN/RL)
L8 6 (L1/SPN OR L1/CPN)

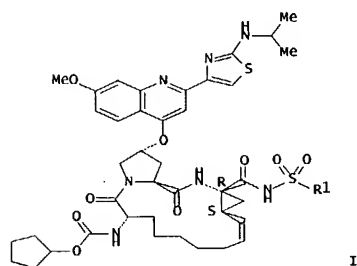
=> FOCUS L8
PROCESSING COMPLETED FOR L8
L9 6 FOCUS L8 1-

=> DIS L9 1- CBIB ABS HTRN
YOU HAVE REQUESTED DATA FROM 6 ANSWERS - CONTINUE? Y/(N):Y

L9 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2004:580783 Document No. 141:261053 Synthesis of BILN 2061, an HCV
NS3

Protease Inhibitor with Proven Antiviral Effect in Humans.
Faucher, Anne-Marie; Bailey, Murray D.; Beaulieu, Pierre L.; Brochu, Christian;
Duceppe, Jean-Simon; Ferland, Jean-Marie; Ghio, Elise; Gorys, Vida;
Halmos, Ted; Kawai, Stephen H.; Poirier, Martin; Simoneau, Bruno;
Tsantrizos, Youla S.; Llinas-Brunet, Montse (Chemistry

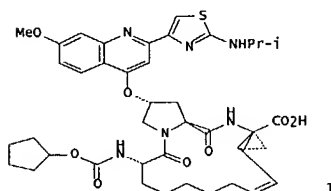
MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU,
SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC,
VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD; RW: AT, BE, BF, BJ, CF, CG,
CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR,
NE, NL, PT, SE, SN, TD, TG, TR. (English). CODEN: PIXXD2.
APPLICATION: WO 2003-CA1604 20031020. PRIORITY: US 2002-PV421414 20021025; US
2002-PV433820 20021216; US 2003-PV442768 20030127.
GI



AB Macrocyclic peptides I [R1 is (un)substituted alkyl, cycloalkyl, alkylcycloalkyl, aryl or heteroaryl] or their pharmaceutically-acceptable salts were prepared as inhibitors of the hepatitis C virus (HCV) NS3 protease. Thus, I (R = Me) was prepared by a multistep sequence involving peptide coupling, olefin metathesis to form the macrocycle and methanesulfonamidation.
IT 300832-84-2P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of macrocyclic peptides active against the hepatitis C virus)

L9 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2004:168624 Document No. 140:350045 Structure-activity study on a novel series of macrocyclic inhibitors of the hepatitis C virus NS3 protease leading to the discovery of BILN 2061. Llinas-Brunet, Montse; Bailey,

Department
Boehringer Ingelheim (Canada) Ltd., Laval, QC, H7S 2G5, Can.).
Organic
Letters, 6(17), 2901-2904 (English) 2004. CODEN: ORLEF7. ISSN:
1523-7060. Publisher: American Chemical Society.
GI



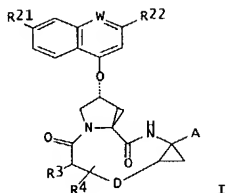
AB The synthesis of BILN 2061 (I), a hepatitis C virus (HCV) NS3 protease inhibitor with proven antiviral effect in humans, was accomplished in a convergent manner from four building blocks. The procedure described here was suitable for the preparation of multigram quantities of BILN 2061 for preclin. pharmacol. evaluation.
IT 300832-84-2P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of peptidyl macrocycle BILN-2061, an HCV NS3 protease inhibitor with proven antiviral effect in humans)

L9 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2004:370958 Document No. 140:357673 Preparation of macrocyclic peptides active against the hepatitis C virus. Llinas-Brunet, Montse; Bailey, Murray D. (Boehringer Ingelheim International G.m.b.h., Germany). PCT Int. Appl. WO 2004037855 A1 20040506, 40 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD,

Murray D.; Bolger, Gordon; Brochu, Christian; Faucher, Anne-Marie; Ferland, Jean Marie; Garneau, Michel; Ghio, Elise; Gorys, Vida; Grand-Maitre, Chantal; Halmos, Ted; Lapeyre-Paquette, Nicole; Liard, Francine; Poirier, Martin; Rheume, Manon; Tsantrizos, Youla S.; Lamarre, Daniel (Departments of Chemistry and Biological Sciences, Boehringer Ingelheim (Canada) Ltd., Laval, QC, H7S 2G5, Can.). Journal of Medicinal Chemistry, 47(7), 1605-1608 (English) 2004. CODEN: JMCMAR. ISSN: 0022-2623. Publisher: American Chemical Society.

AB From the discovery of competitive hexapeptide inhibitors, potent and selective HCV NS3 protease macrocyclic inhibitors have been identified. Structure-activity relationship studies were performed focusing on optimizing the N-terminal carbamate and the aromatic substituent on the (4R)-hydroxyproline moiety. Inhibitors meeting the potency criteria in the cell-based assay and with improved oral bioavailability in rats were identified. BILN 2061 was selected as the best compound, the first NS3 protease inhibitor reported with antiviral activity in man.
IT 300832-84-2P
RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (BILN 2061; structure-activity study on a novel series of macrocyclic inhibitors of the hepatitis C virus NS3 protease leading to the discovery of BILN 2061)

L9 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2003:648255 Document No. 139:197768 Preparation of macrocyclic peptides active against the hepatitis C virus. Tsantrizos, Youla S.; Cameron, Dale R.; Faucher, Anne-Marie; Ghio, Elise; Goudreau, Nathalie; Halmos, Teddy; Llinas-Brunet, Montse (Boehringer Ingelheim (Canada) Ltd., Can.). U.S. US 6608027 B1 20030819, 90 pp., Cont.-in-part of U.S. Ser. No. 542,675, abandoned. (English). CODEN: USXXAM. APPLICATION: US 2001-760946 20010116. PRIORITY: US 1999-PV128011 19990406; US 2000-542675 20000403.
GI



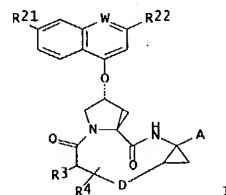
AB Macrocyclic peptides I [W = CH or N; R21 = H, halo, alkyl, cycloalkyl, haloalkyl, alkoxy, cycloalkoxy, hydroxy, or an amino group; R22 = H, halo, alkyl, cycloalkyl, haloalkyl, thioalkyl, alkoxy, cycloalkoxy, alkoxyalkyl, cycloalkyl, aryl or heteroaryl; R3 = hydroxy, NH2, aryl- or heteroaryl amino, NHCOR32, CONHR32, CO2R32, where R32 is alkyl or cycloalkyl; D is a 5 to 10-atom saturated or unsatd. alkylene chain optionally containing one to three heteroatoms independently selected from: O, S, or NH or substituted imino; R4 = H or from one to three substituents at any carbon atom of chain D; A is an amide or carboxylic acid group or a pharmaceutically acceptable salt or ester; two diastereomers may exist at the cyclopropane moiety] were prepared which are active in-vitro and in cellular assays against the NS3 protease of the hepatitis C virus. Thus, macrocyclic peptide I [W = N; R21, R22, R4 = H; A = CO2H; R3CH-D = (S)-(Me3CO2CNH)CH(CH2)3CH:CH(CH2)2-E (syn to acid)] was prepared and showed IC50 > 0.1 µM in the full-length NS3-NS4A heterodimer protein fluorogenic assay.

IT 300832-84-2P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(Preparation of macrocyclic peptides active against the hepatitis C virus)

L9 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS ON STN 2003:511084 Document No. 139:69527 Preparation of macrocyclic compounds as inhibitors of hepatitis C virus. Campbell, Jeffrey Allen; Good, Andrew Charles (Bristol-Myers Squibb Company, USA). PCT Int. Appl. WO 2003053349 A2 20030703, 225 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP,

TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-CA353 20000403. PRIORITY: US 1999-PV128011 19990406.



AB Macrocyclic peptides I [W = CH or N; R21 = H, halo, alkyl, cycloalkyl, haloalkyl, alkoxy, cycloalkoxy, hydroxy, or an amino group; R22 = H, halo, alkyl, cycloalkyl, haloalkyl, thioalkyl, alkoxy, cycloalkoxy, alkoxyalkyl, cycloalkyl, aryl or heteroaryl; R3 = hydroxy, NH2, aryl- or heteroaryl amino, NHCOR32, CONHR32, CO2R32, where R32 is alkyl or cycloalkyl; D is a 5 to 10-atom saturated or unsatd. alkylene chain optionally containing one to three heteroatoms independently selected from: O, S, or NH or substituted imino; R4 = H or from one to three substituents at any carbon atom of chain D; A is an amide or carboxylic acid group or a pharmaceutically acceptable salt or ester; two diastereomers may exist at the cyclopropane moiety] were prepared which are active in-vitro and in cellular assays against the NS3 protease of the hepatitis C virus. Thus, macrocyclic peptide I [W = N; R21, R22, R4 = H; A = CO2H; R3CH-D = (S)-(Me3CO2CNH)CH(CH2)3CH:CH(CH2)2-E (syn to acid)] was prepared and showed IC50 > 0.1 µM in the full-length NS3-NS4A heterodimer protein fluorogenic assay.

IT 300832-84-2P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG, TR. (English). CODEN: PIXXD2. APPLICATION: WO 2002-US39926 20021213. PRIORITY: US 2001-PV344080 20011220; US 2002-PV382103 20020520. GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to macrocyclic compds. I [R1 = (cyclo)alkyl; R2 = H, halo, alkyl, alkoxy, cycloalkoxy, (un)substituted aryl or heterocyclyl; R3 = H, halo, CF3, alkoxy, cycloalkoxy; R4 = NH2 or NHR6, where R6 is alkanoyl, alkylaminocarbonyl, or carbalkoxy; Q is a 3-9 atom (un)saturated alkylene chain optionally containing 1-3 heteroatoms O, S, SO, or SO2], including methods for their synthesis and use in pharmaceutical compns. for therapeutic or prophylactic prevention or treatment of hepatitis C virus (HCV) infection. Thus, 3,13-diazatricyclo[11.3.0.04,6]hexadec-7-ene derivative II was prepared by a multistep procedure and assayed for inhibition of HCV NS3/4A protease (IC50 < 5 µM).

IT 300832-84-2P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (Preparation of macrocyclic compds. as inhibitors of hepatitis C virus)

L9 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS ON STN 2000:725652 Document No. 133:296659 Preparation of macrocyclic peptides active against the hepatitis C virus. Tsantrizos, Youla S.; Cameron, Dale R.; Faucher, Anne-Marie; Ghio, Elise; Goudreau, Nathalie; Halmos, Teddy; Llinas-brunet, Montse (Boehringer Ingelheim (Canada) Ltd., Can.). PCT Int. Appl. WO 2000059929 A1 20001012, 154 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR,

(Uses)
(Preparation of macrocyclic peptides active against the hepatitis C virus)

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STRUCTURE FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6
DICTIONARY FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6

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ACCESSION NUMBER: 2004:252197 HCAPLUS Full-text
DOCUMENT NUMBER: 140:281350
TITLE: Spiro compounds for inhibiting the first-pass effect
INVENTOR(S): Harris, James W.
PATENT ASSIGNEE(S): Bioavailability System, LLC, USA
SOURCE: U.S. Pat. Appl. Publ., 133 pp., Cont.-in-part of U.S.
Ser. No. 793,416.
CODEN: USXXCO
DOCUMENT TYPE: Patent

LANGUAGE: English
FAMILY ACC. NUM. COUNT: 5
PATENT INFORMATION:
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US 6248776 B1 20010619 US 1999-251467
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US 6476066 B1 20021105 US 2001-793416
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20010227 US 1997-56382P P
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DOCUMENT NUMBER: 140:122161
TITLE: An NS3 protease inhibitor with antiviral effects in humans infected with hepatitis C virus
AUTHOR(S): Lamarre, Daniel; Anderson, Paul C.; Bailey, Murray; Beaulieu, Pierre; Bolger, Gordon; Bonneau, Pierre; Boes, Michael; Cameron, Dale R.; Cartier, Mireille; Cordingley, Michael G.; Faucher, Anne-Marie; Goudreau, Nathalie; Kawai, Stephen H.; Kukolj, George; Lagace, Lisette; LaPlante, Steven R.; Narjes, Hans; Poupart, Marc-Andre; Rancourt, Jean; Sentjens, Roel E.; St. George, Roger; Simoneau, Bruno; Steinmann, Gerhard; Thibeault, Diane; Tsantrizos, Youla S.; Weldon, Steven M.; Yong, Chan-Loi; Llinas-Brunet, Montse
CORPORATE SOURCE: Departments of Biological Sciences, Ingelheim (Canada) Ltd, Laval, QC, H7S 2G5, Nature (London, United Kingdom) (2003), 426(6963), 186-189
CODEN: NATUAS; ISSN: 0028-0836
PUBLISHER: Nature Publishing Group

DOCUMENT TYPE: Journal
LANGUAGE: English
REFERENCE COUNT: 30 THERE ARE 30 CITED REFERENCES
AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
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ACCESSION NUMBER: 2000:725652 HCAPLUS Full-text
DOCUMENT NUMBER: 133:296659
TITLE: Preparation of macrocyclic peptides active against the hepatitis C virus
INVENTOR(S): Tsantrizos, Youla S.; Cameron, Dale R.; Faucher, Anne-Marie; Ghio, Elise; Goudreau, Nathalie; Halmos, Teddy; Llinas-brunet, Montse
PATENT ASSIGNEE(S): Boehringer Ingelheim (Canada) Ltd., Can.
SOURCE: PCT Int. Appl., 154 pp., CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

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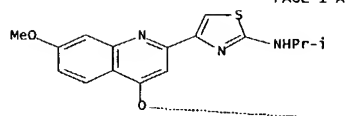
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 WO 2000-CA353 W
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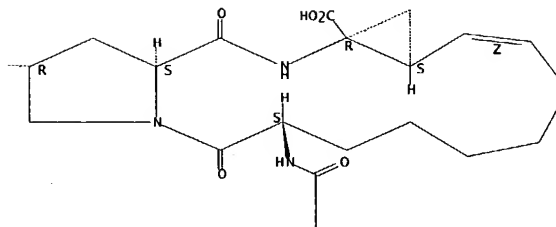
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L14 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
 IT 300832-84-2, BILN 2061
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (spiro compds. for inhibiting the first-pass effect)
 RN 300832-84-2 HCAPLUS
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 carboxylic
 acid, 6-[[[(cyclopentyl)oxy]carbonyl]amino]-
 1,2,3,6,7,8,9,10,11,13a,14,15,16
 16a-tetradecahydro-2-[[7-methoxy-2-[2-[(1-methylethyl)amino]-4-
 thiazolyl]-
 4-quinolinyl]oxy]-5,16-dioxo-, (2R,6S,12Z,13aS,14aR,16aS)- (9CI)
 (CA
 INDEX NAME)

Absolute stereochemistry.
 Double bond geometry as shown.



PAGE 1-B



PAGE 2-B



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2 2061
0 BILN 2061
(BILN(W)2061)
FILE 'MEDLINE'
19 BILN
122 2061

18 2061
0 BILN 2061
(BILN(W)2061)
FILE 'HEALSAFE'
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FILE 'ICONDA'
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FILE 'IFICLS'
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(BILN(W)2061)
FILE 'IFIPAT'
1 BILN
39 2061
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FILE 'IMSDRUGNEWS'
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FILE 'INFODATA'
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0 2061
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FILE 'INIS'
1 BILN
22 2061
0 BILN 2061
(BILN(W)2061)
FILE 'INPADOC'
0 BILN
3 2061
0 BILN 2061
(BILN(W)2061)
FILE 'INSPEC'
0 BILN
73 2061
0 BILN 2061
(BILN(W)2061)
FILE 'INSPHYS'
1 BILN
2 2061
0 BILN 2061
(BILN(W)2061)
FILE 'INVESTEXT'
20 "BILN"
1 "BILNS"
21 "BILN"

13 BILN 2061
(BILN(W)2061)
FILE 'METADEX'
0 BILN
14 2061
0 BILN 2061
(BILN(W)2061)
FILE 'NAPRALERT'
0 "BILN"
2 "2061"
0 BILN 2061
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FILE 'NIOSHITC'
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1 2061
0 BILN 2061
(BILN(W)2061)
FILE 'NLDB'
4 "BILN"
274 "2061"
3 BILN 2061
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FILE 'NTIS'
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27 2061
0 BILN 2061
(BILN(W)2061)
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0 BILN
0 2061
0 BILN 2061
(BILN(W)2061)
FILE 'OCEAN'
0 "BILN"
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FILE 'PAPERCHEM2'
0 BILN
2 2061
0 BILN 2061
(BILN(W)2061)
FILE 'PASCAL'
13 BILN
69 2061
4 BILN 2061
(BILN(W)2061)
FILE 'PATDD'
0 BILN
0 2061
0 BILN 2061
(BILN(W)2061)
FILE 'PATDPA'
0 BILN
5 2061
0 BILN 2061
(BILN(W)2061)

FILE 'PATDPAFULL'
 1 BILN
 210 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'PATOSDE'
 0 BILN
 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'PATOSEP'
 0 BILN
 4 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'PATOSWO'
 0 BILN
 1 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'PCTFULL'
 40 BILN
 2832 2061
 11 BILN 2061
 (BILN(W)2061)
 FILE 'PCTGEN'
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 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'PHARMAML'
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 FILE 'PHIN'
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 FILE 'PIRA'
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 ("BILN"(W)"2061")
 FILE 'POLLUAB'
 0 "BILN"
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 FILE 'PROMT'
 7 "BILN"

538 "2061"
 1 BILN 2061
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 FILE 'RSWB'
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 5 2061
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 (BILN(W)2061)
 FILE 'RUSSCTI'
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 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'SCISEARCH'
 27 BILN
 168 2061
 22 BILN 2061
 (BILN(W)2061)
 FILE 'SIGLE'
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 FILE 'TEMA'
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 12 2061
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 (BILN(W)2061)
 FILE 'TOXCENTER'
 6 BILN
 63 2061
 4 BILN 2061

(BILN(W)2061)
 FILE 'TRIBO'
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 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'TULSA'
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 0 BILN 2061
 (BILN(W)2061)
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 (BILN(W)2061)
 FILE 'UFORDAT'
 0 BILN
 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'ULIDAT'
 0 BILN
 1 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'USPATFULL'
 17 BILN
 4661 2061
 6 BILN 2061
 (BILN(W)2061)
 FILE 'USPAT2'
 0 BILN
 210 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'VETB'
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 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'VETU'
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 3 2061
 0 BILN 2061
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 FILE 'WATER'
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 0 BILN 2061
 (BILN(W)2061)
 FILE 'WELDASEARCH'
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 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'WPIDS'

4 BILN
 70 2061
 2 BILN 2061
 (BILN(W)2061)
 FILE 'WPIFV'
 0 BILN
 0 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'WPINDEX'
 4 BILN
 70 2061
 2 BILN 2061
 (BILN(W)2061)
 FILE 'WSCA'
 0 BILN
 1 2061
 0 BILN 2061
 (BILN(W)2061)
 FILE 'WTEXTILES'
 0 BILN
 8 2061
 0 BILN 2061
 (BILN(W)2061)

L15 QUE BILN 2061

=> d rank
 F1 42 EMBASE
 F2 22 SCISEARCH
 F3 18 BIOSIS
 F4 15 INVESTEXT
 F5 13 MEDLINE
 F6 11 PCTFULL
 F7 10 CAPLUS
 F8 10 DDFU
 F9 10 DRUGU
 F10 8 BIOTECHNO
 F11 6 USPATFULL
 F12 4 LIFESCI
 F13 4 PASCAL
 F14 4 TOXCENTER
 F15 3 ESRIOBASE
 F16 3 IMSDRUGNEWS
 F17 3 NLDB
 F18 2 ADISCTI
 F19 2 BIOENG
 F20 2 CRNB
 F21 2 COMPENDEX
 F22 2 PHIN
 F23 2 WPIDS
 F24 2 WPINDEX
 F25 1 BABS
 F26 1 CIN
 F27 1 EMBAL
 F28 1 IFIPAT
 F29 1 PROMT

=> index f1-f29
 COST IN U.S. DOLLARS
 TOTAL
 SESSION
 FULL ESTIMATED COST
 94.22
 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
 TOTAL
 SESSION
 CA SUBSCRIBER PRICE
 4.90
 INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
 CAPLUS, DDFU,
 DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
 ESBIOBASE,
 IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
 WPIDS,
 WPINDEX, BABS, CIN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20
 ON 26 OCT 2004

29 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view
 search error messages that display as 0* with SET DETAIL OFF.

=> d his 114-

(FILE 'HCAPLUS' ENTERED AT 14:32:46 ON 26 OCT 2004)
 L14 3 L10 AND PD<20030327

FILE 'HOME' ENTERED AT 14:39:45 ON 26 OCT 2004

INDEX 'IMOBILITY, ZMOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
 AGRICOLA,
 ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
 BABS,
 BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
 BIOTECHARS,
 BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...' ENTERED AT
 15:24:55 ON
 26 OCT 2004

SEA BILN 2061

2 FILE ADISCTI
 1 FILE BABS
 2 FILE BIOENG
 18 FILE BIOSIS
 8 FILE BIOTECHNO
 10 FILE CAPLUS
 2 FILE CBNB
 1 FILE CIN

20 "BILN"
 1 "BILNS"
 21 "BILN"
 ("BILN" OR "BILNS")
 1864 "2061"
 15 BILN 2061
 ("BILN"(w)"2061")
 34452 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'MEDLINE'
 19 BILN
 122 2061
 13 BILN 2061
 (BILN(w)2061)
 122732 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'PCTFULL'
 40 BILN
 2832 2061
 11 BILN 2061
 (BILN(w)2061)
 127299 CRYSTAL?
 9 L15 AND CRYSTAL?
 FILE 'CAPLUS'
 20 BILN
 382 2061
 10 BILN 2061
 (BILN(w)2061)
 1605797 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'DDFU'
 13 BILN
 21 2061
 10 BILN 2061
 (BILN(w)2061)
 7961 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'DRUGU'
 13 BILN
 36 2061
 10 BILN 2061
 (BILN(w)2061)
 12117 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'BIOTECHNO'
 12 BILN
 43 2061
 8 BILN 2061
 (BILN(w)2061)
 27086 CRYSTAL?
 3 L15 AND CRYSTAL?
 FILE 'USPATFULL'
 17 BILN
 4661 2061
 6 BILN 2061
 (BILN(w)2061)
 570101 CRYSTAL?

2 FILE COMPENDEX
 10 FILE DDFU
 10 FILE DRUGU
 1 FILE EMBAL
 42 FILE EMBASE
 3 FILE ESBIOBASE
 1 FILE IFIPAT
 3 FILE IMSDRUGNEWS
 15 FILE INVESTEXT
 4 FILE LIFESCI
 13 FILE MEDLINE
 3 FILE NLDB
 4 FILE PASCAL
 11 FILE PCTFULL
 2 FILE PHIN
 1 FILE PROMT
 22 FILE SCISEARCH
 4 FILE TOXCENTER
 6 FILE USPATFULL
 2 FILE WPIDS
 2 FILE WPINDEX
 L15 QUE BILN 2061

INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
 CAPLUS,
 DDFU, DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
 ESBIOBASE,
 IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
 WPIDS,
 WPINDEX, BABS, CIN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20 ON
 26 OCT
 2004

=> 115 and crystal?

FILE 'EMBASE'
 48 "BILN"
 124 "2061"
 42 BILN 2061
 ("BILN"(w)"2061")
 97097 CRYSTAL?
 4 L15 AND CRYSTAL?
 FILE 'SCISEARCH'
 27 BILN
 168 2061
 22 BILN 2061
 (BILN(w)2061)
 689047 CRYSTAL?
 3 L15 AND CRYSTAL?
 FILE 'BIOSIS'
 27 BILN
 129 2061
 18 BILN 2061
 (BILN(w)2061)
 111782 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'INVESTEXT'

6 L15 AND CRYSTAL?
 FILE 'LIFESCI'
 5 "BILN"
 28 "2061"
 4 BILN 2061
 ("BILN"(w)"2061")
 27296 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'PASCAL'
 13 BILN
 69 2061
 4 BILN 2061
 (BILN(w)2061)
 517682 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'TOXCENTER'
 6 BILN
 63 2061
 4 BILN 2061
 (BILN(w)2061)
 66181 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'ESBIOBASE'
 3 BILN
 49 2061
 3 BILN 2061
 (BILN(w)2061)
 41699 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'IMSDRUGNEWS'
 3 "BILN"
 5 "2061"
 3 BILN 2061
 ("BILN"(w)"2061")
 118 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'NLDB'
 4 "BILN"
 274 "2061"
 3 BILN 2061
 ("BILN"(w)"2061")
 32830 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'ADISCTI'
 3 BILN
 21 2061
 2 BILN 2061
 (BILN(w)2061)
 463 CRYSTAL?
 0 L15 AND CRYSTAL?
 FILE 'BIOENG'
 2 BILN
 7 2061
 2 BILN 2061
 (BILN(w)2061)
 7912 CRYSTAL?
 0 L15 AND CRYSTAL?

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FILE 'CBNB'
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  15 2061
  2 BILN 2061
    (BILN(W)2061)
  5823 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'COMPENDEX'
  4 BILN
  22 2061
  2 BILN 2061
    (BILN(W)2061)
  443469 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'PHIN'
  3 "BILN"
  35 "2061"
  2 BILN 2061
    ("BILN"(W)"2061")
  1251 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'WPIDS'
  4 BILN
  70 2061
  2 BILN 2061
    (BILN(W)2061)
  362675 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'WPINDEX'
  4 BILN
  70 2061
  2 BILN 2061
    (BILN(W)2061)
  362675 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'BABS'
  1 BILN
  24 2061
  1 BILN 2061
    (BILN(W)2061)
  91052 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'CIN'
  1 "BILN"
  12 "2061"
  1 BILN 2061
    ("BILN"(W)"2061")
  6321 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'EMBAL'
  1 BILN
  4 2061
  1 BILN 2061
    (BILN(W)2061)
  1192 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'IFIPAT'

```

```

=> L16
L18      25 L16

=> dup rem
ENTER L# LIST OR (END):L18
PROCESSING COMPLETED FOR L18
L19      21 DUP REM L18 (4 DUPLICATES REMOVED)

```

```

=> L19 and pd<20030327
  1 FILES SEARCHED...
  3 FILES SEARCHED...
L20      2 L19 AND PD<20030327

```

```

=> d l20 1-2 ibib hitstr abs kwic
'HITSTR' IS NOT A VALID FORMAT FOR FILE 'EMBASE'

```

The following are valid formats:

The default display format is BIB.

```

ABS ----- AB
ALL ----- AN, DN, TI, AU, CS, SO, PUI, CY, DT, FS, LA, SL, AB,
           CT, RN, CN, NP, CO, GEN
BIB ----- AN, DN, TI, AU, CS, SO, PUI, CY, DT, FS, LA, SL
CBIB ----- Compressed bibliographic data
DALL ----- ALL, delimited for post-processing
IABS ----- ABS, with a text label
IALL ----- ALL, indented with text labels
IBIB ----- BIB, indented with text labels
IND ----- CT, RN, CN, NP, CO, GEN
TRIAL ----- TI, CT, RN, CN, NP, CO, GEN
(SAM, TRI)
HIT ----- All fields containing hit terms
HITIND ----- IND
KWIC ----- All hit terms plus 20 words on either side
OCC ----- List of display fields containing hit terms
           and number of occurrences in each field

```

Hit terms will be highlighted in all displayable fields.

To display a particular field or fields, enter the display field codes. For a list of display field codes, enter 'HELP DFIELDS' at an arrow prompt (=>). Examples of formats include: 'BIB'; 'AB'; 'SO,SI'. You may specify the format fields in any order, and the information will be displayed in the same order as the format specification.

The same formats (except for HIT, HITIND, KWIC, and OCC) may be used with the DISPLAY ACC command to display the record for a specified Accession Number.

ENTER DISPLAY FORMAT (BIB):ia11

L20 ANSWER 1 OF 2 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.
ON STN

```

  1 BILN
  39 2061
  1 BILN 2061
    (BILN(W)2061)
  146324 CRYSTAL?
  0 L15 AND CRYSTAL?
FILE 'PROMT'
  7 "BILN"
  538 "2061"
  1 BILN 2061
    ("BILN"(W)"2061")
  95992 CRYSTAL?
  0 L15 AND CRYSTAL?

```

L16 QUE L15 AND CRYSTAL?

```

=> d rank
F1      9 PCTFULL
F2      6 USPATFULL
F3      4 EMBASE
F4      3 SCISEARCH
F5      3 BIOTECHNO

```

```

=> file f3-5,f1,f2
COST IN U.S. DOLLARS
TOTAL

```

SINCE FILE

ENTRY

```

SESSION
FULL ESTIMATED COST
95.93

```

1.71

```

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL

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SINCE FILE

ENTRY

```

SESSION
CA SUBSCRIBER PRICE
4.90

```

0.00

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FILE 'USPATFULL' ENTERED AT 15:27:48 ON 26 OCT 2004
CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

```

=> L16, dup rem
L17      0 L16, DUP REM

```

ACCESSION NUMBER: 2003468113 EMBASE Full-text
TITLE: Current therapy and new molecular approaches to antiviral

AUTHOR: Hugle T.; Cerny A.
CORPORATE SOURCE: Dr. A. Cerny, Clinica Medica, Ospedale Civico, CH-6903

SOURCE: Lugano, Switzerland. andreas.cerny@bluewin.ch
Reviews in Medical Virology, (2003) 13/6 (361-371).

Refs: 79
ISSN: 1052-9276 CODEN: RMVIEW
COUNTRY: United Kingdom
DOCUMENT TYPE: Journal; General Review
FILE SEGMENT: 004 Microbiology
030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
039 Pharmacy

LANGUAGE: English
SUMMARY LANGUAGE: English

ABSTRACT:
Current therapeutic options for hepatitis C are limited, especially for genotype 1. For genotypes 2 and 3, pegylated interferon in combination with ribavirin, can lead to a sustained virological response in up to 80% of patients. Unfortunately, adverse effects of IFN and ribavirin are a major problem and the list of contraindications for HCV therapy is long, including decompensated cirrhosis of the liver and psychiatric disorders. Therefore, alternative therapeutic approaches are needed. New delivery options for IFN and ribavirin are aimed at optimising efficiency and reducing adverse effects. Recent progress in the molecular virology of HCV has identified new targets for antiviral intervention. Inhibition of HCV gene expression and replication as well as immunotherapeutic concepts aimed at enhancing the cellular immune response against HCV are being explored. Solution of the crystal structures of HCV key enzymes led to the design of specific inhibitors including compounds active against the well characterised NS3 serine protease and RNA-dependent RNA polymerase which are currently in the early phase clinical investigation. New strategies for inhibiting HCV gene expression include the use of antisense oligodeoxynucleotides and ribozymes. Immunomodulation by agents such as inosine monophosphate dehydrogenase inhibitors, thymosin-alpha 1, histamine or amantadine are being

studied in combination with IFN and/or ribavirin. Immunotherapeutic vaccination with recombinant HCV E1 protein improved host immunity against HCV and thus seems to be a promising new option. Copyright .COPYRG. 2003 John Wiley & Sons, Ltd.

CONTROLLED TERM: Medical Descriptors:
 *hepatitis C: DT, drug therapy
 *hepatitis C: ET, etiology
 *hepatitis C: PC, prevention
 *infection prevention
 virus gene
 genotype
 drug response
 drug contraindication
 drug delivery system
 side effect: SI, side effect
 gene expression
 drug targeting
 immunotherapy
 enzyme structure
 crystal structure
 drug design
 drug activity
 antiviral activity
 protein targeting
 immunomodulation
 vaccination
 Hepatitis C virus
 immune response
 cellular immunity
 hemolytic anemia: SI, side effect
 mental disease: SI, side effect
 flu like syndrome: SI, side effect
 leukopenia: SI, side effect
 thrombocytopenia: SI, side effect
 teratogenicity
 virus replication
 drug hypersensitivity: SI, side effect
 rash: SI, side effect
 human
 nonhuman
 clinical trial
 review
 Drug Descriptors:
 alpha interferon: AE, adverse drug reaction
 alpha interferon: CT, clinical trial
 alpha interferon: CB, drug combination
 alpha interferon: DT, drug therapy
 alpha interferon: TO, drug toxicity
 alpha interferon: PR, pharmacokinetics
 alpha interferon: PD, pharmacology
 alpha interferon: SC, subcutaneous drug
 ribavirin: AE, adverse drug reaction

administration

trial

combination

therapy

pharmacology

CAS REGISTRY NO.:
 37205-61-1;

198821-22-6,

CHEMICAL NAME:

COMPANY NAME:

NABI;

ribozyme: CT, clinical trial
 ribozyme: DT, drug therapy
 ribozyme: TO, drug toxicity
 ribozyme: PD, pharmacology
 hepatocyte: AE, adverse drug reaction
 hepatocyte: CT, clinical trial
 hepatocyte: DT, drug therapy
 hepatocyte: TO, drug toxicity
 hepatocyte: PD, pharmacology
 antisense oligodeoxynucleotide: CT, clinical
 antisense oligodeoxynucleotide: DT, drug therapy
 antisense oligodeoxynucleotide: PD, pharmacology
 Isis 14803: CT, clinical trial
 Isis 14803: DT, drug therapy
 Isis 14803: PD, pharmacology
 RNA derivative: DV, drug development
 RNA derivative: DT, drug therapy
 RNA derivative: PD, pharmacology
 small interfering rna: DV, drug development
 small interfering rna: DT, drug therapy
 small interfering rna: PD, pharmacology
 monoclonal antibody: DT, drug therapy
 monoclonal antibody: PD, pharmacology
 xtl 002: DT, drug therapy
 xtl 002: PD, pharmacology
 cicavir: DT, drug therapy
 cicavir: PD, pharmacology
 immunomodulating agent: CB, drug combination
 immunomodulating agent: DT, drug therapy
 thymosin alpha: CT, clinical trial
 thymosin alpha: CB, drug combination
 thymosin alpha: DO, drug dose
 thymosin alpha: DT, drug therapy
 thymosin alpha: PD, pharmacology
 inosinate dehydrogenase inhibitor: CB, drug
 inosinate dehydrogenase inhibitor: DT, drug
 inosinate dehydrogenase inhibitor: PD,
 merimepodib: CT, clinical trial
 merimepodib: CB, drug combination
 merimepodib: DT, drug therapy
 merimepodib: PD, pharmacology
 unindexed drug
 unclassified drug
 (ribavirin) 36791-04-5; (proteinase inhibitor)
 (thymosin alpha) 69521-94-4; (merimepodib)
 198821-38-4
 (1) vx 950; (2) Jtk 003; Biln 2061; Isis 14803;
 xtl 002
 (1) Vertex; (2) Akros; Ribozyme Pharmaceuticals;
 Sciclone; Regenerx; Maxim

ribavirin: CT, clinical trial
 ribavirin: CB, drug combination
 ribavirin: CM, drug comparison
 ribavirin: DT, drug therapy
 ribavirin: PK, pharmacokinetics
 ribavirin: PD, pharmacology
 ribavirin: PO, oral drug administration
 albumin conjugate: PR, pharmacokinetics
 liposome: PR, pharmacokinetics
 polyaminoacid: PR, pharmacokinetics
 polyaminoacid: PO, oral drug administration
 ribavirin derivative: AE, adverse drug reaction
 ribavirin derivative: CT, clinical trial
 ribavirin derivative: CB, drug combination
 ribavirin derivative: CM, drug comparison
 ribavirin derivative: DT, drug therapy
 ribavirin derivative: PD, pharmacology
 viramidine: AE, adverse drug reaction
 viramidine: CT, clinical trial
 viramidine: CB, drug combination
 viramidine: CM, drug comparison
 viramidine: DT, drug therapy
 viramidine: PD, pharmacology
 levovirin: AE, adverse drug reaction
 levovirin: CT, clinical trial
 levovirin: CM, drug comparison
 levovirin: DT, drug therapy
 levovirin: PD, pharmacology
 proteinase inhibitor: AE, adverse drug reaction
 proteinase inhibitor: CT, clinical trial
 proteinase inhibitor: DO, drug dose
 proteinase inhibitor: DT, drug therapy
 proteinase inhibitor: PK, pharmacokinetics
 proteinase inhibitor: PD, pharmacology
 proteinase inhibitor: PO, oral drug
 administration
 biln 2061: AE, adverse drug reaction
 biln 2061: CT, clinical trial
 biln 2061: DO, drug dose
 biln 2061: DT, drug therapy
 biln 2061: PK, pharmacokinetics
 biln 2061: PD, pharmacology
 biln 2061: PO, oral drug administration
 vx 950: DT, drug therapy
 vx 950: PD, pharmacology
 virus protein
 protein NS5B
 RNA directed DNA polymerase inhibitor: CT,
 clinical trial
 RNA directed DNA polymerase inhibitor: DT, drug
 therapy
 RNA directed DNA polymerase inhibitor: PD,
 pharmacology
 jtk 003: CT, clinical trial
 jtk 003: DT, drug therapy
 jtk 003: PD, pharmacology
 ribozyme: AE, adverse drug reaction

L20 ANSWER 2 OF 2 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS
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 on STN
 ACCESSION NUMBER: 2003195244 EMBASE Full-text
 TITLE: Hepatitis C virus therapies: Current treatments,
 targets
 and future perspectives.
 AUTHOR: Walker M.P.; Appleby T.C.; Zhong W.; Lau J.Y.N.;
 Hong Z.
 CORPORATE SOURCE: Z. Hong, Ribapharm Inc., Hyland Avenue, Costa
 Mesa, CA,
 SOURCE: United States. zhihong@ribapharm.com
 Antiviral Chemistry and Chemotherapy, (2003) 14/1
 (1-21).
 Refs: 208
 ISSN: 0956-3202 CODEN: ACCHEH
 COUNTRY: United Kingdom
 DOCUMENT TYPE: Journal; General Review
 FILE SEGMENT: 004 Microbiology
 030 Pharmacology
 037 Drug Literature Index
 038 Adverse Reactions Titles
 048 Gastroenterology
 LANGUAGE: English
 SUMMARY LANGUAGE: English
 ABSTRACT:
 Chronic hepatitis C virus (HCV) infection is the cause of an emerging
 global
 epidemic of chronic liver disease. Current combination therapies are
 at best
 80% efficacious and are often poorly tolerated. Strategies to improve
 the
 therapeutic response include the development of novel interferons,
 nucleoside
 analogues with reduced haemolysis compared with ribavirin and inosine
 5'-monophosphate dehydrogenase inhibitors. Compounds in preclinical
 or early
 clinical trials include small molecules that inhibit virus-specific
 enzymes
 (such as the serine proteases, RNA polymerase and helicase) or
 interfere with
 translation (including antisense molecules, iRNA and ribozymes).
 Advances in
 understanding HCV replication, obtaining a sub-genomic replicon and
 contriving
 potential small animal models, in addition to solving
 crystallographic
 structures for the replication enzymes, have improved prospects for
 developing
 novel therapies. This review summarizes current and evolving
 treatments for
 chronic hepatitis C infection. In addition, progress in HCV targets
 and drug
 discovery tools valuable in the search for novel anti-HCV agents is
 detailed.

CONTROLLED TERM:

Medical Descriptors:
 *hepatitis C: DT, drug therapy
 *hepatitis C: EP, epidemiology
 *hepatitis C: ET, etiology
 *chronic liver disease: ET, etiology
 drug efficacy
 drug tolerance
 hemolytic anemia: SI, side effect
 side effect: SI, side effect
 alanine aminotransferase blood level
 virus replication
 replicon
 crystal structure
 RNA translation
 untranslated region
 internal ribosome entry site
 monotherapy
 virus load
 treatment outcome
 treatment indication
 immunomodulation
 drug safety
 treatment failure
 chimpanzee
 transgenic mouse
 Hepatitis GB virus B
 IC 50
 structure activity relation
 drug structure
 virus assembly
 human
 nonhuman
 clinical trial
 review
 priority journal
 Drug Descriptors:
 *antivirus agent: AE, adverse drug reaction
 *antivirus agent: CT, clinical trial
 *antivirus agent: AN, drug analysis
 *antivirus agent: CB, drug combination
 *antivirus agent: CM, drug comparison
 *antivirus agent: DV, drug development
 *antivirus agent: DO, drug dose
 *antivirus agent: DT, drug therapy
 *antivirus agent: PD, pharmacology
 *antivirus agent: IV, intravenous drug
 *antivirus agent: SC, subcutaneous drug
 alpha interferon: AE, adverse drug reaction
 alpha interferon: CB, drug combination
 alpha interferon: CM, drug comparison
 alpha interferon: DO, drug dose
 alpha interferon: DT, drug therapy
 alpha interferon: PD, pharmacology
 nucleoside derivative: AN, drug analysis
 nucleoside derivative: CM, drug comparison

administration

administration

administration

levovirin: DV, drug development
 levovirin: DO, drug dose
 levovirin: DT, drug therapy
 levovirin: PD, pharmacology
 viramidine: CT, clinical trial
 viramidine: AN, drug analysis
 viramidine: CM, drug comparison
 viramidine: DV, drug development
 viramidine: DO, drug dose
 viramidine: DT, drug therapy
 viramidine: PD, pharmacology
 merimepodib: CT, clinical trial
 merimepodib: AN, drug analysis
 merimepodib: CB, drug combination
 merimepodib: CM, drug comparison
 merimepodib: DV, drug development
 merimepodib: DT, drug therapy
 merimepodib: PD, pharmacology
 thymosin alpha1: CT, clinical trial
 thymosin alpha1: AN, drug analysis
 thymosin alpha1: CB, drug combination
 thymosin alpha1: DV, drug development
 thymosin alpha1: DO, drug dose
 thymosin alpha1: DT, drug therapy
 thymosin alpha1: PD, pharmacology
 thymosin alpha1: SC, subcutaneous drug

amantadine: CT, clinical trial
 amantadine: AN, drug analysis
 amantadine: CB, drug combination
 amantadine: CM, drug comparison
 amantadine: DV, drug development
 amantadine: PD, pharmacology
 recombinant interleukin 12: CT, clinical trial
 recombinant interleukin 12: AN, drug analysis
 recombinant interleukin 12: CB, drug combination
 recombinant interleukin 12: CM, drug comparison
 recombinant interleukin 12: DV, drug development
 recombinant interleukin 12: DO, drug dose
 recombinant interleukin 12: DT, drug therapy
 recombinant interleukin 12: PD, pharmacology
 histamine: CT, clinical trial
 histamine: AN, drug analysis
 histamine: CB, drug combination
 histamine: DV, drug development
 histamine: DT, drug therapy
 histamine: PD, pharmacology
 gamma interferon: CT, clinical trial
 gamma interferon: AN, drug analysis
 gamma interferon: CB, drug combination
 gamma interferon: DV, drug development
 gamma interferon: DT, drug therapy
 gamma interferon: PD, pharmacology
 proteinase inhibitor: CT, clinical trial
 proteinase inhibitor: DO, drug dose
 proteinase inhibitor: PD, pharmacology
 proteinase inhibitor: PO, oral drug

comparison

therapy

pharmacology

comparison

drug

comparison

drug

administration

nucleoside derivative: DV, drug development
 nucleoside derivative: PR, pharmaceuticals
 nucleoside derivative: PD, pharmacology
 ribavirin: AE, adverse drug reaction
 ribavirin: CT, clinical trial
 ribavirin: CB, drug combination
 ribavirin: CM, drug comparison
 ribavirin: DO, drug dose
 ribavirin: DT, drug therapy
 ribavirin: PD, pharmacology
 inosinate dehydrogenase inhibitor: CM, drug
 inosinate dehydrogenase inhibitor: DT, drug
 inosinate dehydrogenase inhibitor: PD,
 serine proteinase: EC, endogenous compound
 RNA polymerase: EC, endogenous compound
 helicase: EC, endogenous compound
 ribozyme: EC, endogenous compound
 recombinant alpha2a interferon: CM, drug
 recombinant alpha2a interferon: DO, drug dose
 recombinant alpha2a interferon: DT, drug therapy
 recombinant alpha2a interferon: PD, pharmacology
 recombinant alpha2a interferon: SC, subcutaneous
 administration
 recombinant alpha2b interferon: CM, drug
 recombinant alpha2b interferon: DO, drug dose
 recombinant alpha2b interferon: DT, drug therapy
 recombinant alpha2b interferon: PD, pharmacology
 recombinant alpha2b interferon: SC, subcutaneous
 administration
 consensus interferon: CM, drug comparison
 consensus interferon: DO, drug dose
 consensus interferon: DT, drug therapy
 consensus interferon: PD, pharmacology
 consensus interferon: SC, subcutaneous drug
 peginterferon alpha2b: CT, clinical trial
 peginterferon alpha2b: CB, drug combination
 peginterferon alpha2b: CM, drug comparison
 peginterferon alpha2b: DO, drug dose
 peginterferon alpha2b: DT, drug therapy
 peginterferon alpha2b: PD, pharmacology
 peginterferon alpha2a: CT, clinical trial
 peginterferon alpha2a: CB, drug combination
 peginterferon alpha2a: CM, drug comparison
 peginterferon alpha2a: DO, drug dose
 peginterferon alpha2a: DT, drug therapy
 peginterferon alpha2a: PD, pharmacology
 levovirin: CT, clinical trial
 levovirin: AN, drug analysis
 levovirin: CM, drug comparison

administration

benzocyclohepten

tetrahydropyran

CAS REGISTRY NO.:

6;

(peginterferon

198153-51-4;

alpha1)

(histamine)

[4 [[6,7

tetrahydropyran 4

CHEMICAL NAME:

779; Amd

biln 2061: CT, clinical trial
 biln 2061: DO, drug dose
 biln 2061: PD, pharmacology
 biln 2061: PO, oral drug administration
 peptide derivative: AN, drug analysis
 peptide derivative: DV, drug development
 peptide derivative: PD, pharmacology
 peptide alpha keto acid: AN, drug analysis
 peptide alpha keto acid: DV, drug development
 peptide alpha keto acid: PD, pharmacology
 pyrrolidine derivative: AN, drug analysis
 pyrrolidine derivative: DV, drug development
 pyrrolidine derivative: PD, pharmacology
 pyrrolidine 5,5 lactam: AN, drug analysis
 pyrrolidine 5,5 lactam: DV, drug development
 pyrrolidine 5,5 lactam: PD, pharmacology
 iddb3: DV, drug development
 iddb3: PD, pharmacology
 unindexed drug
 unclassified drug
 isis 14803
 gw 3112
 gw 2549
 gw 0569
 n [4 [[6,7 dihydro 2 (4 methylphenyl) 5h
 8 yl]carbonyl]amino]benzyl] n,n dimethyl 2h
 4 aminium chloride
 1,1' [1,4 phenylenebis(methylene)]bis(1,4,8,11
 tetraazacyclotetradecane)
 (ribavirin) 36791-04-5; (serine proteinase)
 (RNA polymerase) 9014-24-8; (helicase) 42613-29-
 (recombinant alpha2b interferon) 98530-12-2;
 alpha2b) 215647-85-1; (peginterferon alpha2a)
 (merimepodib) 198821-22-6, 198821-38-4; (thymosin
 69521-94-4; (amantadine) 665-66-7, 768-94-5;
 51-45-6, 56-92-8, 93443-21-1; (gamma interferon)
 82115-62-6; (proteinase inhibitor) 37205-61-1; (n
 dihydro 2 (4 methylphenyl) 5h benzocyclohepten 8
 yl]carbonyl]amino]benzyl] n,n dimethyl 2h
 aminium chloride) 229005-80-5; (1,1' [1,4
 phenylenebis(methylene)]bis(1,4,8,11
 tetraazacyclotetradecane)) 155148-31-5
 (1) vx 497; (2) Ceplene; (3) Biln 2061; (4) Isis
 14803; Zadaxin; Gw 3112; Gw 2549; Gw 0569; Tak
 3100; iddb3

COMPANY NAME: (1) Vertex (United States); (2) Maxim; (3)
Boehringer
Merck
Kingdom); Bristol
States);
Ingelheim; (4) Isis (United States); Ribapharm;
(United States); Glaxo SmithKline (United
Myers Squibb (United States); Celera (United
Viropharma; Japanese tobacco; IRBM

=> d his

(FILE 'HOME' ENTERED AT 14:24:22 ON 26 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE,
AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS,
BIOTECHDS,
BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN,
CONFSCI, CROPR,
CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 14:25:23 ON
26 OCT 2004

L1 FILE 'REGISTRY' ENTERED AT 14:26:32 ON 26 OCT 2004
1 S 300832-84-2/RN
SET SMA OFF
DEL SEL Y
SEL RN
SET SMA LOGIN

INDEX 'BEILSTEIN, GMELIN' ENTERED AT 14:27:43 ON 26 OCT 2004
SEA E1 AND OPTICAL?/FA

L2 QUE 300832-84-2/BI AND OPTICAL?/FA

L3 FILE 'CAPLUS' ENTERED AT 14:28:02 ON 26 OCT 2004
0 S L1/BMF

L4 FILE 'CAPLUS' ENTERED AT 14:28:18 ON 26 OCT 2004
0 S L1/BPN

L5 FILE 'CAPLUS' ENTERED AT 14:28:28 ON 26 OCT 2004
0 S L1/IMF

L6 FILE 'CAPLUS' ENTERED AT 14:28:41 ON 26 OCT 2004
1 S L1/PEP

L7 FILE 'CAPLUS' ENTERED AT 14:29:30 ON 26 OCT 2004
0 S L1/PUR

L8 FILE 'CAPLUS' ENTERED AT 14:29:48 ON 26 OCT 2004
6 S (L1/SPN OR L1/CPN)
L9 6 FOCUS L8 1-

FILE 'REGISTRY' ENTERED AT 14:31:45 ON 26 OCT 2004

IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
WPIDS,
WPINDEX, BABS, CIN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20 ON
26 OCT 2004

SEA L15 AND CRYSTAL?

L16 4 FILE EMBASE
3 FILE SCISEARCH
9 FILE PCTFULL
3 FILE BIOTECHNO
6 FILE USPATFULL
QUE L15 AND CRYSTAL?

FILE 'EMBASE, SCISEARCH, BIOTECHNO, PCTFULL, USPATFULL' ENTERED

AT 15:27:48 ON 26 OCT 2004

L17 0 L16, DUP REM
L18 25 L16
L19 21 DUP REM L18 (4 DUPLICATES REMOVED)
L20 2 L19 AND PD<20030327

=> stnindex

ENTER FILE OR CLUSTER NAMES (NONE):all
FILE 'ENCOMPLIT' ACCESS NOT AUTHORIZED
FILE 'ENCOMPLIT2' ACCESS NOT AUTHORIZED
FILE 'ENCOMPPAT' ACCESS NOT AUTHORIZED
FILE 'ENCOMPPAT2' ACCESS NOT AUTHORIZED

COST IN U.S. DOLLARS
TOTAL

SINCE FILE

ENTRY

SESSION
FULL ESTIMATED COST
112.45

16.52

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL

SINCE FILE

ENTRY

SESSION
CA SUBSCRIBER PRICE
4.90

0.00

INDEX 'IMOBILITY, ZMOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA,
ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS,
BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS,
BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...'
ENTERED AT 15:32:46 ON 26 OCT 2004

143 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view
search error messages that display as 0* with SET DETAIL OFF.

FILE 'HCAPLUS' ENTERED AT 14:32:46 ON 26 OCT 2004
L10 15 S L1
L11 0 L10 AND CRYSTAL
L12 0 L10 AND CRYSTAL?
L13 0 L10 AND ALCOHOL
L14 3 L10 AND PD<20030327

FILE 'HOME' ENTERED AT 14:39:45 ON 26 OCT 2004

INDEX 'IMOBILITY, ZMOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA,
ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS,
BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS,
BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...' ENTERED AT
15:24:55 ON
26 OCT 2004

SEA BILN 2061

2 FILE ADISCTI
1 FILE BABS
2 FILE BIOENG
18 FILE BIOSIS
8 FILE BIOTECHNO
10 FILE CAPLUS
2 FILE CBNB
1 FILE CIN
2 FILE COMPENDEX
10 FILE DDFU
10 FILE DRUGU
1 FILE EMBAL
42 FILE EMBASE
3 FILE ESBIODBASE
1 FILE IFIPAT
3 FILE IMSDRUGNEWS
15 FILE INVESTEXT
4 FILE LIFESCI
13 FILE MEDLINE
3 FILE NLDB
4 FILE PASCAL
11 FILE PCTFULL
2 FILE PHIN
1 FILE PROMT
22 FILE SCISEARCH
4 FILE TOXCENTER
6 FILE USPATFULL
2 FILE WPIDS
2 FILE WPINDEX
L15 QUE BILN 2061

INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
CAPLUS,
DDFU, DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
ESBIODBASE,

=> ciluprevir/cn
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FILE 'ZMOBILITY'
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-----User Break-----
'CN' IS NOT A VALID FIELD CODE
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FILE 'BIOSIS'
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SEARCH ENDED BY USER
FILE 'BIOTECHABS'
SEARCH ENDED BY USER

=> ciluprevir/
FILE 'IMOBILITY'
'CILUPREVIR/' IS NOT A VALID FIELD CODE
For a list of field codes for the current file, enter "HELP SFIELDS"
at an arrow prompt (=>).

=> ciluprevir
FILE 'IMOBILITY'
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 FILE 'RAPRA' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'RSWB' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'RUSSCI' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'SCISEARCH' 2 CILUPREVIR
 FILE 'SIGLE' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'SOLIDSTATE' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'SOLIS' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'SYNTHLINE' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'TEMA' 0 CILUPREVIR
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 FILE 'TEXTILETECH' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'TOXCENTER' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'TRIBO' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'TULSA' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'TULSA2' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'UFORDAT' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'ULIDAT' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'USPATFULL' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'USPAT2' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'VETB' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'VETU' 0 CILUPREVIR

FILE 'NIOSTHC' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'NLDB' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'NTIS' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'NUTRACEUT' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'OCEAN' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PAPERCHEM2' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PASCAL' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PATDD' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PATDPA' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PATDPAFULL' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PATOSDE' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PATOSEP' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PATOSWO' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PCTFULL' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PCTGEN' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PHARMAML' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PHIC' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PHIN' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PIRA' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'POLLUAB' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'PROMT'

0 CILUPREVIR
 FILE 'WATER' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'WELDASEARCH' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'WPIDS' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'WPIFV' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'WPINDE' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'WSCA' 0 CILUPREVIR
 0 CILUPREVIR
 FILE 'WTEXTILES' 0 CILUPREVIR
 0 CILUPREVIR

L21 QUE CILUPREVIR

=> d rank		
F1	3	EMBASE
F2	2	ESBIODASE
F3	2	MEDLINE
F4	2	SCISEARCH
F5	1	DDFU
F6	1	DRUGU

=> fil f1-6		
COST IN U.S. DOLLARS		SINCE FILE
TOTAL		ENTRY

SESSION		
FULL ESTIMATED COST		5.70
118.15		

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)		SINCE FILE
TOTAL		ENTRY

SESSION		
CA SUBSCRIBER PRICE		0.00
4.90		

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FILE 'DDFU' ACCESS NOT AUTHORIZED

FILE 'DRUGU' ENTERED AT 15:39:01 ON 26 OCT 2004
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=> s 121
L22 10 L21
=> 122 and pd<20030327
1 FILES SEARCHED...
2 FILES SEARCHED...
'20030327' NOT A VALID FIELD CODE
'20030327' NOT A VALID FIELD CODE
L23 1 L22 AND PD<20030327

=> d 123

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on STN
AN 2004039472 EMBASE Full-text
TI Gateways to Clinical Trials.
AU Bayes M.; Rabasseda X.; Prous J.R.
CS M. Bayes, Prous Science, S.A., P.O. Box 540, 08080 Barcelona, Spain.
mbayes@prous.com
SO Methods and Findings in Experimental and Clinical Pharmacology,
(
2003) 25/10 (831-855).
Refs: 145
ISSN: 0379-0355 CODEN: MFEPDX
CY Spain
DT Journal; General Review
FS 030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
LA English
SL English

=> FIL STNGUIDE
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
126.52
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL
SESSION
CA SUBSCRIBER PRICE
SINCE FILE
ENTRY
8.37
SINCE FILE
ENTRY
0.00

exenatide, exisulind, ezetimib; Febuxostat; Gallium maltolate, ganirelix acetate, garenoxacin mesylate, gefitinib; H11, HuMax; IL-15, IDD-1, IGIV-C, imatinib mesylate, ISIS-14803, ITF-1697, ivabradine hydrochloride; KRN-5500; L-365260, levetiracetam, levosimendan, licoferone, linezolid, LJP-1082, lopinavir, lumiracoxib; MCC-478, melatonin, morphine hydrochloride, morphine-6-glucuronide, moxidectin; N-Acetylcarnosine, natalizumab, NM-702, NNC-05-1869, NSC-703940; Ocinaon OM-89, omalizumab, omeprazole/sodium bicarbonate, OPC-28326, ospemifene; PEG-filgrastim peginterferon alfa-2a, pegsunercept, pirfenidone, pralmorelin, pregabalin; Recombinant glucagon-like peptide-1 (7-36) amide, repifermin, RSD-1235; S-8184, selodnoson, sodium dichloroacetate, suberanolhydroxamic acid; TAS-102, terfenadine, teriparatide, tipranavir troxycitabine; Ximelagatran; YM-337. .COPYRG. 2003 Prous Science. All rights reserved.

CT Medical Descriptors:
*drug monitoring
drug indication
drug efficacy
drug safety
side effect: SI, side effect
patient compliance
drug tolerability
liver toxicity: SI, side effect
bleeding: SI, side effect
disease exacerbation
systemic lupus erythematosus: SI, side effect
digestive system ulcer: SI, side effect
neutropenia: SI, side effect
teratogenicity: SI, side effect
human
clinical trial
review
Drug Descriptors:
abetimus: CT, clinical trial
abetimus: IV, intravenous drug administration
adalimumab: AE, adverse drug reaction
adalimumab: CT, clinical trial
linezolid: CT, clinical trial
alemtuzumab: CT, clinical trial
ivabradine: CT, clinical trial
ivabradine: IV, intravenous drug administration
recombinant interleukin 1 receptor blocking agent: CT, clinical trial
recombinant interleukin 1 receptor blocking agent: IA,
intraarterial drug
administration
glucagon like peptide 1: CT, clinical trial
glucagon like peptide 1: SC, subcutaneous drug administration
astemizole: CT, clinical trial
atazanavir: CT, clinical trial
bosentan: CT, clinical trial
botulinum toxin B: CT, clinical trial
caspofungin: CT, clinical trial
ciclesonide: CT, clinical trial
cilomilast: CT, clinical trial

4.90

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AND TECHNOLOGY CORPORATION, AND FACHINFORMATIONSZENTRUM KARLSRUHE

FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Oct 22, 2004 (20041022/UP).

=> 123 all
MISSING OPERATOR L23 ALL
The search profile that was entered contains terms or
nested terms that are not separated by a logical operator.
=> d 123 all
YOU HAVE REQUESTED DATA FROM FILE 'EMBASE' - CONTINUE? (Y)/N:y

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AN 2004039472 EMBASE Full-text
TI Gateways to Clinical Trials.
AU Bayes M.; Rabasseda X.; Prous J.R.
CS M. Bayes, Prous Science, S.A., P.O. Box 540, 08080 Barcelona, Spain.
mbayes@prous.com
SO Methods and Findings in Experimental and Clinical Pharmacology,
(
2003) 25/10 (831-855).
Refs: 145
ISSN: 0379-0355 CODEN: MFEPDX
CY Spain
DT Journal; General Review
FS 030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
LA English
SL English
AB Gateways to Clinical Trials is a guide to the most recent clinical trials in current literature and congresses. The data in the following tables has been retrieved from the Clinical Studies Knowledge Area of Prous Science Integrity®, the drug discovery and development portal, <http://integrity.prous.com>. This issue focuses on the following selection of drugs: Abetimus sodium, adalimumab, alefacept, alemtuzumab, almotriptan, AMG-0007, anakinra, anti-CTLA-4 Mab, L-arginine hydrochloride, arxoxifene hydrochloride, astemizole, atazanavir sulfate, atizumab; Belimumab, BG-9928, binodenoson, bosentan, botulinum toxin type B, bovine lactoferrin, BufferGel; Caspofungin acetate, ciclesonide, cilomilast, ciluprevir, clofarabine, CVT-3146; Darbepoetin alfa, desloratadine, diflomotecan, doripenem, dronedarone hydrochloride, drotrecogin alfa (activated), DT388-GM-CSF, duloxetine hydrochloride, E-5564, efalizumab, enfuvirtide, esomeprazole magnesium, estradiol acetate, ETC-642,

efalizumab: CT, clinical trial
imatinib: CT, clinical trial
terfenadine: CT, clinical trial
tipranavir: CT, clinical trial
tipranavir: PO, oral drug administration
ximelagatran: CT, clinical trial
ximelagatran: PO, oral drug administration
ym 337: CT, clinical trial
moxidectin: CT, clinical trial
estradiol: CT, clinical trial
novel erythropoiesis stimulating protein: CT, clinical trial
novel erythropoiesis stimulating protein: IV, intravenous drug administration
novel erythropoiesis stimulating protein: SC, subcutaneous drug administration
desloratadine: CT, clinical trial
desloratadine: PO, oral drug administration
diflomotecan: CT, clinical trial
diflomotecan: IV, intravenous drug administration
diflomotecan: PO, oral drug administration
morphine: CT, clinical trial
etiracetam: CT, clinical trial
doripenem: CT, clinical trial
duloxetine: CT, clinical trial
unindexed drug
RN (abetimus) 167362-48-3, 169147-32-4; (adalimumab) 331731-18-1; (linezolid) 165800-03-3; (alemtuzumab) 216503-57-0; (ivabradine) 148849-67-6, 148870-80-8, 155974-00-8; (glucagon like peptide 1) 89750-14-1; (astemizole) 68844-77-9; (atazanavir) 198904-31-3; (bosentan) 147536-97-8, 157212-55-0; (caspofungin) 189768-38-5; (ciclesonide) 126544-47-6; (cilomilast) 153259-65-5; (efalizumab) 214745-43-4; (imatinib) 152459-95-5, 220127-57-1; (terfenadine) 50679-08-8; (tipranavir) 174484-41-4; (ximelagatran) 192939-46-1, 260790-58-7; (moxidectin) 113507-06-5; (estradiol) 50-28-2; (desloratadine) 100643-71-8; (diflomotecan) 220997-97-7; (morphine) 52-26-6, 57-27-2; (etiracetam) 102767-28-2, 33996-58-6; (doripenem) 148016-81-3; (duloxetine) 116539-59-4, 136434-34-9
CN Ym 337

=> DIS HIST

(FILE 'HOME' ENTERED AT 14:24:22 ON 26 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 14:25:23 ON

26 OCT 2004

L1 FILE 'REGISTRY' ENTERED AT 14:26:32 ON 26 OCT 2004
1 S 300832-84-2/RN
SET SMA OFF
DEL SEL Y
SEL RN
SET SMA LOGIN
INDEX 'BEILSTEIN, GELIN' ENTERED AT 14:27:43 ON 26 OCT 2004
SEA E1 AND OPTICAL7/FA
L2 QUE 300832-84-2/BI AND OPTICAL7/FA
FILE 'CAPLUS' ENTERED AT 14:28:02 ON 26 OCT 2004
0 S L1/BMF
L4 FILE 'CAPLUS' ENTERED AT 14:28:18 ON 26 OCT 2004
0 S L1/BPN
L5 FILE 'CAPLUS' ENTERED AT 14:28:28 ON 26 OCT 2004
0 S L1/IMF
L6 FILE 'CAPLUS' ENTERED AT 14:28:41 ON 26 OCT 2004
1 S L1/PEP
L7 FILE 'CAPLUS' ENTERED AT 14:29:30 ON 26 OCT 2004
0 S L1/PUR
L8 FILE 'CAPLUS' ENTERED AT 14:29:48 ON 26 OCT 2004
6 S (L1/SPN OR L1/CPN)
L9 6 FOCUS L8 1-
FILE 'REGISTRY' ENTERED AT 14:31:45 ON 26 OCT 2004
FILE 'HCAPLUS' ENTERED AT 14:32:46 ON 26 OCT 2004
L10 15 S L1
L11 0 L10 AND CRYSTAL
L12 0 L10 AND CRYSTAL?
L13 0 L10 AND ALCOHOL
L14 3 L10 AND PD<20030327
FILE 'HOME' ENTERED AT 14:39:45 ON 26 OCT 2004
INDEX 'IMOBILITY, 2MOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA,
ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS,
BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS,
BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...' ENTERED AT
15:24:55 ON
26 OCT 2004
SEA BILN 2061
2 FILE ADISCTI

INDEX 'IMOBILITY, 2MOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA,
ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS,
BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS,
BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...' ENTERED AT
15:32:46 ON
26 OCT 2004
SEA CILUPREVIR/CN
0* FILE IMOBILITY
0* FILE 2MOBILITY
0* FILE ADISCTI
0* FILE AEROSPACE
0* FILE ALUMINIUM
0* FILE ANTE
0* FILE APOLLIT
0* FILE AQUALINE
0* FILE AQUASCI
0* FILE BABS
0* FILE BIBLIODATA
0* FILE BIOCOMMERCE
0* FILE BIOENG
SEA CILUPREVIR/
0* FILE IMOBILITY
SEA CILUPREVIR
1 FILE DDFU
1 FILE DRUGU
3 FILE EMBASE
2 FILE ESBIODBASE
2 FILE MEDLINE
2 FILE SCISEARCH
L21 QUE CILUPREVIR

FILE 'EMBASE, ESBIODBASE, MEDLINE, SCISEARCH, DRUGU' ENTERED AT
15:39:01
ON 26 OCT 2004
L22 10 S L21
L23 1 L22 AND PD<20030327
FILE 'STNGUIDE' ENTERED AT 15:40:30 ON 26 OCT 2004
FILE 'EMBASE' ENTERED AT 15:41:46 ON 26 OCT 2004
FILE 'STNGUIDE' ENTERED AT 15:41:46 ON 26 OCT 2004

=>

---Logging off of STN---

1 FILE BABS
2 FILE BIOENG
18 FILE BIOSIS
8 FILE BIOTECHNO
10 FILE CAPLUS
2 FILE CBNB
1 FILE CIN
2 FILE COMPENDEX
10 FILE DDFU
10 FILE DRUGU
1 FILE EMBAL
42 FILE EMBASE
3 FILE ESBIODBASE
1 FILE IFIPAT
3 FILE IMSDRUGNEWS
15 FILE INVESTEXT
4 FILE LIFESCI
13 FILE MEDLINE
3 FILE NLDB
4 FILE PASCAL
11 FILE PCTFULL
2 FILE PHIN
1 FILE PROMT
22 FILE SCISEARCH
4 FILE TOXCENTER
6 FILE USPATFULL
2 FILE WPIDS
2 FILE WPINDEX
L15 QUE BILN 2061
INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
CAPLUS,
DDFU, DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
ESBIODBASE,
IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
WPIDS,
WPINDEX, BABS, CIN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20 ON
26 OCT
2004
SEA L15 AND CRYSTAL?
4 FILE EMBASE
3 FILE SCISEARCH
9 FILE PCTFULL
3 FILE BIOTECHNO
6 FILE USPATFULL
L16 QUE L15 AND CRYSTAL?
FILE 'EMBASE, SCISEARCH, BIOTECHNO, PCTFULL, USPATFULL' ENTERED
AT
15:27:48 ON 26 OCT 2004
L17 0 L16, DUP REM
L18 25 L16
L19 21 DUP REM L18 (4 DUPLICATES REMOVED)
L20 2 L19 AND PD<20030327

=>
Executing the logoff script...

=> LOG Y

COST IN U.S. DOLLARS	SINCE FILE
TOTAL	ENTRY
SESSION	0.12
FULL ESTIMATED COST	
130.58	
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE
TOTAL	ENTRY
SESSION	0.00
CA SUBSCRIBER PRICE	
4.90	

STN INTERNATIONAL LOGOFF AT 15:42:43 ON 26 OCT 2004

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:sssptal653adk

PASSWORD:
TERMINAL (ENTER 1, 2, 3, OR ?):2

***** Welcome to STN International *****

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America
NEWS 2 "Ask CAS" for self-help around the clock
NEWS 3 Jul 12 BEILSTEIN enhanced with new display and select
options,
resulting in a closer connection to BABS
NEWS 4 AUG 02 IFIPAT/IFTUDB/IFICDB reloaded with new search and
display fields
NEWS 5 AUG 02 Caplus and CA patent records enhanced with European
and Japan
Patent Office Classifications
NEWS 6 AUG 02 The Analysis Edition of STN Express with Discover!
(Version 7.01 for Windows) now available
NEWS 7 AUG 27 BIOCOMMERCE: changes and enhancements to content
coverage
NEWS 8 AUG 27 BIOTECHABS/BIOTECHDS: Two new display fields added
for legal status data from INPADOC
NEWS 9 SEP 01 INPADOC: New family current-awareness alert (SDI)
available
NEWS 10 SEP 01 New pricing for the Save Answers for SciFinder
Wizard within
STN Express with Discover!
NEWS 11 SEP 01 New display format, HITSTR, available in
WPIDS/WPINDEX/WPIX
NEWS 12 SEP 27 STANDARDS will no longer be available on STN
NEWS 13 SEP 27 SWETSCAN will no longer be available on STN
NEWS EXPRESS JULY 30 CURRENT WINDOWS VERSION IS V7.01, CURRENT
MACINTOSH VERSION IS V6.0c(ENG) AND V6.0jc(JP),
AND CURRENT DISCOVER FILE IS DATED 11 AUGUST 2004
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NEWS INTER Direct Dial and Telecommunication Network Access to
STN
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Enter NEWS followed by the item number or name to see news on that

Please note that search-term pricing does apply when
conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for
details.

Experimental and calculated property data are now available. For more
information enter HELP PROP at an arrow prompt in the file or refer
to the file summary sheet on the web at:
<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> S 300832-84-2/RN
L1 1 300832-84-2/RN

=> DEL SEL Y

=> SEL RN
E1 THROUGH E1 ASSIGNED

=> INDEX BEILSTEIN, GMELIN
COST IN U.S. DOLLARS
TOTAL

SINCE FILE
ENTRY

SESSION
FULL ESTIMATED COST 0.92
2.48

INDEX 'BEILSTEIN, GMELIN' ENTERED AT 14:27:43 ON 26 OCT 2004

2 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view
search error messages that display as 0* with SET DETAIL OFF.

=> S E1 AND OPTICAL?/FA
FILE 'BEILSTEIN'
0 300832-84-2/BI
(300832-84-2/RN)
753186 OPTICAL?/FA
0 300832-84-2/BI AND OPTICAL?/FA
FILE 'GMELIN'
0 300832-84-2/BI
(300832-84-2/RN)
12899 OPTICAL?/FA
0 300832-84-2/BI AND OPTICAL?/FA

L2 QUE 300832-84-2/BI AND OPTICAL?/FA

=> DIS RANK
NO F-NUMBERS HAD GREATER THAN ZERO HITS

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL

SINCE FILE
ENTRY

specific topic.

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FILE 'HOME' ENTERED AT 14:24:22 ON 26 OCT 2004

=> index biosci
FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED
COST IN U.S. DOLLARS
TOTAL

SINCE FILE
ENTRY

SESSION
FULL ESTIMATED COST 0.42
0.42

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE,
AQUALINE,
AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS,
BIOTECHDS,
BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN,
CONFSCI, CROPB,
CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 14:25:23 ON
26 OCT 2004

75 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view
search error messages that display as 0* with SET DETAIL OFF.

=> fil reg
COST IN U.S. DOLLARS
TOTAL

SINCE FILE
ENTRY

SESSION
FULL ESTIMATED COST 1.14
1.56

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STRUCTURE FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6
DICTIONARY FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

SESSION
FULL ESTIMATED COST 0.57
3.05

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FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate
substance identification.

=> S L1/BMF
15 L1
52155 BMF/RL
L3 0 L1/BMF
(L1 (L) BMF/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL

SINCE FILE
ENTRY

SESSION
FULL ESTIMATED COST 2.26
5.31

FILE 'CAPLUS' ENTERED AT 14:28:18 ON 26 OCT 2004
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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/BPN
15 L1
102210 BPN/RL
L4 0 L1/BPN
(L1 (L) BPN/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
7.57
SINCE FILE
ENTRY
2.26

FILE 'CAPLUS' ENTERED AT 14:28:28 ON 26 OCT 2004
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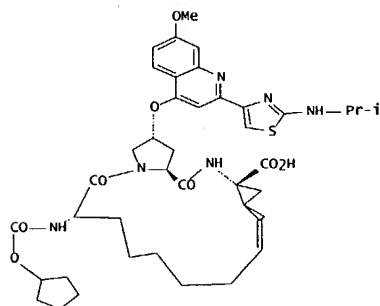
FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/IMF
15 L1
390491 IMF/RL
L5 0 L1/IMF
(L1 (L) IMF/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
7.57
SINCE FILE
ENTRY
2.26

KZ, MD; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG, TR.
(English). CODEN: PIXXD2. APPLICATION: WO 2003-US30402
20030925.
PRIORITY: US 2002-PV414940 20020930; US 2002-PV421904 20021029;
US 2002-PV433834 20021216; US 2003-PV443662 20030130.
GI



AB Disclosed are oral pharmaceutical compns., kits and methods of treating and preventing Hepatitis C Viral (HCV) infections wherein Compound (I), a potent inhibitor of HCV serine protease, or a pharmaceutically acceptable salt thereof, is administered in a selected dosage range. Also disclosed are the use of I or a pharmaceutically acceptable salt thereof, as a control substance for validating an HCV replication assay and also as a control substance for determining the relative effectiveness of one or more substances, alone or in combination, to inhibit the replication of HCV.

IT 300832-84-2
RL: PAC (Pharmacological activity); PEP (Physical, engineering or chemical process); PYP (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (potent inhibitor of HCV serine protease)

SESSION
FULL ESTIMATED COST
9.83

2.26

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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/PEP
15 L1
1687596 PEP/RL
L6 1 L1/PEP
(L1 (L) PEP/RL)

=> DIS L6 1 CBIB ABS HITRN

L6 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2004 ACS ON STN
2004:310970 Document No. 140:327091 Potent inhibitor of HCV serine protease.
Chen, Shirley; Nehmiz, Gerhard; Croenlein, Jens Oliver; Steinmann, Gerhard; Gunn, Jocelyn Abella; Costa, Phuong Do (Boehringer Ingelheim International G.m.b.H., Germany). PCT Int. Appl. WO 2004030670 A1
20040415, 42 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG,

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
14.95
SINCE FILE
ENTRY
5.12
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL
SESSION
CA SUBSCRIBER PRICE
0.70
SINCE FILE
ENTRY
-0.70

FILE 'CAPLUS' ENTERED AT 14:29:30 ON 26 OCT 2004
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FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S L1/PUR
15 L1
201134 PUR/RL
L7 0 L1/PUR
(L1 (L) PUR/RL)

=> FIL CAPLUS
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
17.21
SINCE FILE
ENTRY
2.26
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL
SESSION
ENTRY

CA SUBSCRIBER PRICE
0.70

0.00

FILE 'CAPLUS' ENTERED AT 14:29:48 ON 26 OCT 2004
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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> S (L1/SPN OR L1/CPN)

15 L1
1660968 SPN/RL
6 L1/SPN
(L1 (L) SPN/RL)

15 L1
1155 CPN/RL
0 L1/CPN
(L1 (L) CPN/RL)

L8 6 (L1/SPN OR L1/CPN)

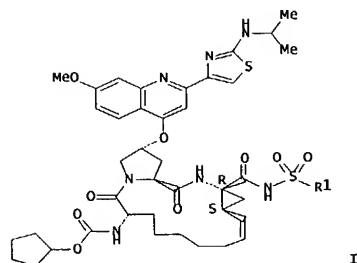
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PROCESSING COMPLETED FOR L8
L9 6 FOCUS L8 1-

=> DIS L9 1- CBIB ABS HITRN
YOU HAVE REQUESTED DATA FROM 6 ANSWERS - CONTINUE? Y/(N):Y

L9 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2004:580783 Document No. 141:261053 Synthesis of BILN 2061, an HCV
NS3

Protease Inhibitor with Proven Antiviral Effect in Humans.
Faucher, Anne-Marie; Bailey, Murray D.; Beaulieu, Pierre L.; Brochu, Christian; Duceppe, Jean-Simon; Ferland, Jean-Marie; Ghiro, Elise; Gorys, Vida; Halmos, Ted; Kawai, Stephen H.; Poirier, Martin; Simoneau, Bruno; Tsantrizos, Youla S.; Llinas-Brunet, Montse (Chemistry

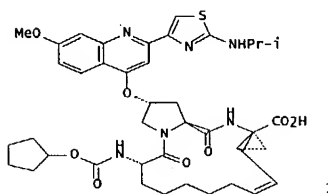
MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG, TR. (English). CODEN: PIXXD2.
APPLICATION: WO 2003-CA1604 20031020. PRIORITY: US 2002-PV421414 20021025; US 2002-PV433820 20021216; US 2003-PV442768 20030127.
GI



AB Macrocyclic peptides I [R1 is (un)substituted alkyl, cycloalkyl, alkylcycloalkyl, aryl or heteroaryl] or their pharmaceutically-acceptable salts were prepared as inhibitors of the hepatitis C virus (HCV) NS3 protease. Thus, I (R = Me) was prepared by a multistep sequence involving peptide coupling, olefin metathesis to form the macrocycle and methanesulfonamidation.
IT 300832-84-2P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)
(preparation of macrocyclic peptides active against the hepatitis C virus)

L9 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2004:168624 Document No. 140:350045 Structure-activity study on a novel series of macrocyclic inhibitors of the hepatitis C virus NS3 protease leading to the discovery of BILN 2061. Llinas-Brunet, Montse; Bailey,

Department, Boehringer Ingelheim (Canada) Ltd., Laval, QC, H7S 2G5, Can.).
Organic Letters, 6(17), 2901-2904 (English) 2004. CODEN: ORLEF7. ISSN: 1523-7060. Publisher: American Chemical Society.
GI



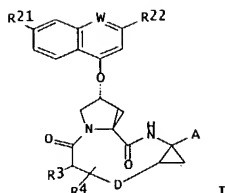
AB The synthesis of BILN 2061 (I), a hepatitis C virus (HCV) NS3 protease inhibitor with proven antiviral effect in humans, was accomplished in a convergent manner from four building blocks. The procedure described here was suitable for the preparation of multigram quantities of BILN 2061 for preclin. pharmacol. evaluation.
IT 300832-84-2P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(preparation of peptidyl macrocycle BILN-2061, an HCV NS3 protease inhibitor with proven antiviral effect in humans)

L9 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2004:370958 Document No. 140:357673 Preparation of macrocyclic peptides active against the hepatitis C virus. Llinas-Brunet, Montse; Bailey, Murray D. (Boehringer Ingelheim International G.m.b.h., Germany). PCT Int. Appl. WO 2004037855 A1 20040506, 40 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD,

Murray D.; Bolger, Gordon; Brochu, Christian; Faucher, Anne-Marie; Ferland, Jean Marie; Garneau, Michel; Ghiro, Elise; Gorys, Vida; Grand-Maitre, Chantal; Halmos, Ted; Lapeyre-Paquette, Nicole; Liard, Francine; Poirier, Martin; Rheume, Manon; Tsantrizos, Youla S.; Lamarre, Daniel (Departments of Chemistry and Biological Sciences, Boehringer Ingelheim (Canada) Ltd., Laval, QC, H7S 2G5, Can.). Journal of Medicinal Chemistry, 47(7), 1605-1608 (English) 2004. CODEN: JMCMAR. ISSN: 0022-2623. Publisher: American Chemical Society.

AB From the discovery of competitive hexapeptide inhibitors, potent and selective HCV NS3 protease macrocyclic inhibitors have been identified. Structure-activity relationship studies were performed focusing on optimizing the N-terminal carbamate and the aromatic substituent on the (4R)-hydroxyproline moiety. Inhibitors meeting the potency criteria in the cell-based assay and with improved oral bioavailability in rats were identified. BILN 2061 was selected as the best compound, the first NS3 protease inhibitor reported with antiviral activity in man.
IT 300832-84-2P
RL: PAC (Pharmacological activity); PRP (Properties); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation) (BILN 2061; structure-activity study on a novel series of macrocyclic inhibitors of the hepatitis C virus NS3 protease leading to the discovery of BILN 2061)

L9 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
2003:648255 Document No. 139:197768 Preparation of macrocyclic peptides active against the hepatitis C virus. Tsantrizos, Youla S.; Cameron, Dale R.; Faucher, Anne-Marie; Ghiro, Elise; Goudreau, Nathalie; Halmos, Teddy; Llinas-Brunet, Montse (Boehringer Ingelheim (Canada) Ltd., Can.). U.S. 6608027 B1 20030819, 90 pp., Cont.-in-part of U.S. Ser. No. 542,675, abandoned. (English). CODEN: USXXAM. APPLICATION: US 2001-760946 20010116. PRIORITY: US 1999-PV128011 19990406; US 2000-542675 20000403.
GI



KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG, TR. (English). CODEN: PIXXD2. APPLICATION: WO 2002-0539926. 20021213. PRIORITY: US 2001-PV344080 20011220; US 2002-PV382103 20020520. GI

* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB The invention relates to macrocyclic compds. I [R1 = (cyclo)alkyl; R2 = H, halo, alkyl, alkoxy, cycloalkoxy, (un)substituted aryl or heterocyclyl; R3 = H, halo, CF3, alkoxy, cycloalkoxy; R4 = NH2 or NHR6, where R6 is alkanoyl, alkylaminocarbonyl, or carbalkoxy; Q is a 3-9 atom (un)saturated alkylene chain optionally containing 1-3 heteroatoms O, S, SO, or SO2], including methods for their synthesis and use in pharmaceutical compns. for therapeutic or prophylactic prevention or treatment of hepatitis C virus (HCV) infection. Thus, 3,13-diazatetracyclo[11.3.0.0.4,6]hexadec-7-ene derivative II was prepared by a multistep procedure and assayed for inhibition of HCV NS3/4A protease (IC50 < 5 µM).

IT 300832-84-2P
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (preparation of macrocyclic compds. as inhibitors of hepatitis C virus)

L9 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN 2000:725652 Document No. 133:296659 Preparation of macrocyclic peptides

active against the hepatitis C virus. Tzantrizos, Youla S.; Cameron, Dale R.; Faucher, Anne-marie; Ghio, Elise; Goudreau, Nathalie; Halmos, Teddy; Llinas-brunet, Montse (Boehringer Ingelheim (Canada) Ltd., Can.). PCT Int. Appl. WO 2000059929 A1 20001012, 154 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR,

(Uses)
(preparation of macrocyclic peptides active against the hepatitis C virus)

=> FIL REGISTRY
COST IN U.S. DOLLARS
TOTAL
SESSION
FULL ESTIMATED COST
42.78
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
TOTAL
SESSION
CA SUBSCRIBER PRICE
4.90
SINCE FILE
ENTRY
25.57
SINCE FILE
ENTRY
-4.20

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STRUCTURE FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6
DICTIONARY FILE UPDATES: 25 OCT 2004 HIGHEST RN 769101-30-6

TSCA INFORMATION NOW CURRENT THROUGH MAY 21, 2004

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<http://www.cas.org/ONLINE/DBSS/registryss.html>

=> d his

(FILE 'HOME' ENTERED AT 14:24:22 ON 26 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROBP, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 14:25:23 ON 26 OCT 2004

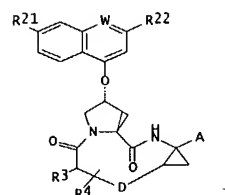
AB Macrocyclic peptides I [W = CH or N; R21 = H, halo, alkyl, cycloalkyl, haloalkyl, alkoxy, cycloalkoxy, hydroxy, or an amino group; R22 = H, halo, alkyl, cycloalkyl, haloalkyl, thioalkyl, alkoxy, cycloalkoxy, alkoxyalkyl, cycloalkyl, aryl or heteroaryl; R3 = hydroxy, NH2, aryl- or heteroaryl amino, NHCOR32, CONHR32, CO2R32, where R32 is alkyl or cycloalkyl; D is a 5 to 10-atom saturated or unsatd. alkylene chain optionally containing one to three heteroatoms independently selected from: O, S, or NH or substituted imino; R4 = H or from one to three substituents at any carbon atom of chain D; A is an amide or carboxylic acid group or a pharmaceutically acceptable salt or ester; two diastereomers may exist at the cyclopropane moiety] were prepared which are active in-vitro and in cellular assays against the NS3 protease of the hepatitis C virus. Thus, macrocyclic peptide I [W = N; R21, R22, R4 = H; A = CO2H; R3CH-D = (S)-(Me3CO2CNH)CH(CH2)3CH:CH(CH2)2-E (syn to acid)] was prepared and showed IC50 > 0.1 µM in the full-length NS3-NS4A heterodimer protein fluorogenic assay.

IT 300832-84-2P
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of macrocyclic peptides active against the hepatitis C virus)

L9 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN 2003:511084 Document No. 139:69527 Preparation of macrocyclic compounds as inhibitors of hepatitis C virus. Campbell, Jeffrey Allen; Good, Andrew Charles (Bristol-Myers Squibb Company, USA). PCT Int. Appl. WO 2003053349 A2 20030703, 225 pp. DESIGNATED STATES: W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP,

TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM; RW: AT, BE, BF, BJ, CF, CG, CH, CI, CM, CY, DE, DK, ES, FI, FR, GA, GB, GR, IE, IT, LU, MC, ML, MR, NE, NL, PT, SE, SN, TD, TG. (English). CODEN: PIXXD2. APPLICATION: WO 2000-CA353 20000403. PRIORITY: US 1999-PV128011 19990406. GI



AB Macrocyclic peptides I [W = CH or N; R21 = H, halo, alkyl, cycloalkyl, haloalkyl, alkoxy, cycloalkoxy, hydroxy, or an amino group; R22 = H, halo, alkyl, cycloalkyl, haloalkyl, thioalkyl, alkoxy, cycloalkoxy, alkoxyalkyl, cycloalkyl, aryl or heteroaryl; R3 = hydroxy, NH2, aryl- or heteroaryl amino, NHCOR32, CONHR32, CO2R32, where R32 is alkyl or cycloalkyl; D is a 5 to 10-atom saturated or unsatd. alkylene chain optionally containing one to three heteroatoms independently selected from: O, S, or NH or substituted imino; R4 = H or from one to three substituents at any carbon atom of chain D; A is an amide or carboxylic acid group or a pharmaceutically acceptable salt or ester; two diastereomers may exist at the cyclopropane moiety] were prepared which are active in-vitro and in cellular assays against the NS3 protease of the hepatitis C virus. Thus, macrocyclic peptide I [W = N; R21, R22, R4 = H; A = CO2H; R3CH-D = (S)-(Me3CO2CNH)CH(CH2)3CH:CH(CH2)2-E (syn to acid)] was prepared and showed IC50 > 0.1 µM in the full-length NS3-NS4A heterodimer protein fluorogenic assay.

IT 300832-84-2P
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

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SEL RN
SET SMA LOGIN

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SEA E1 AND OPTICAL?/FA

L2 QUE 300832-84-2/BI AND OPTICAL?/FA

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0 S L1/BMF

L4 FILE 'CAPLUS' ENTERED AT 14:28:18 ON 26 OCT 2004
0 S L1/BPN

L5 FILE 'CAPLUS' ENTERED AT 14:28:28 ON 26 OCT 2004
0 S L1/IMF

L6 FILE 'CAPLUS' ENTERED AT 14:28:41 ON 26 OCT 2004
1 S L1/PEP

L7 FILE 'CAPLUS' ENTERED AT 14:29:30 ON 26 OCT 2004
0 S L1/PUR

L8 FILE 'CAPLUS' ENTERED AT 14:29:48 ON 26 OCT 2004
6 S (L1/SPN OR L1/CPN)
L9 6 FOCUS L8 1-

FILE 'REGISTRY' ENTERED AT 14:31:45 ON 26 OCT 2004

=> file hcaplus
COST IN U.S. DOLLARS SINCE FILE
TOTAL ENTRY

SESSION
FULL ESTIMATED COST 0.84
43.62

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE
TOTAL ENTRY

SESSION
CA SUBSCRIBER PRICE 0.00
4.90

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FILE COVERS 1907 - 26 Oct 2004 VOL 141 ISS 18
FILE LAST UPDATED: 25 Oct 2004 (20041025/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s l1
L10 15 L1

=> l10 and crystal
1117482 CRYSTAL
600101 CRYSTALS
1385437 CRYSTAL
(CRYSTAL OR CRYSTALS)
L11 0 L10 AND CRYSTAL

=> l10 and crystal?
1605797 CRYSTAL?
L12 0 L10 AND CRYSTAL?

=> l10 and alcohol
217243 ALCOHOL
149677 ALCOHOLS
339514 ALCOHOL
(ALCOHOL OR ALCOHOLS)
L13 0 L10 AND ALCOHOL

=> l10 and pd<20030327
23271868 PD<20030327
(PD<20030327)
L14 3 L10 AND PD<20030327

=> d l14 1-3 ibib

L14 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2004:252197 HCAPLUS Full-text
DOCUMENT NUMBER: 140:281350
TITLE: Spiro compounds for inhibiting the first-pass effect
INVENTOR(S): Harris, James W.
PATENT ASSIGNEE(S): Bioavailability System, LLC, USA
SOURCE: U.S. Pat. Appl. Publ., 133 pp., Cont.-in-part of U.S.
Ser. No. 793,416.
CODEN: USXXCO
DOCUMENT TYPE: Patent

LANGUAGE: English
FAMILY ACC. NUM. COUNT: 5
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO.
DATE -----

US 2004058982 A1 20040325 US 2003-422848
20030425
US 6248776 B1 20010619 US 1999-251467
19990217 <--
US 6476066 B1 20021105 US 2001-793416
20010227 <--
PRIORITY APPLN. INFO.: US 1999-251467 A3
19990217 US 2001-793416 A2
20010227 US 1997-56382P P
19970826 US 1997-997259 A2
19971223
OTHER SOURCE(S): MARPAT 140:281350

L14 ANSWER 2 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2003:886572 HCAPLUS Full-text
DOCUMENT NUMBER: 140:122161
TITLE: An NS3 protease inhibitor with antiviral effects in humans infected with hepatitis C virus
AUTHOR(S): Lamarre, Daniel; Anderson, Paul C.; Bailey, Murray; Beaulieu, Pierre; Bolger, Gordon; Bonneau, Pierre; Boes, Michael; Cameron, Dale R.; Cartier, Mireille; Cordingley, Michael G.; Faucher, Anne-Marie; Goudreau, Nathalie; Kawai, Stephen H.; Kukolj, George; Lagace, Lisette; LaPlante, Steven R.; Narjes, Hans; Poupart, Marc-Andre; Rancourt, Jean; Sentjens, Roel E.; St. George, Roger; Simoneau, Bruno; Steinmann, Gerhard; Thibeault, Diane; Tsantrizos, Youla S.; Weldon, Steven
CORPORATE SOURCE: M.; Yong, Chan-Loi; Llinas-Brunet, Montse
Boehringer Departments of Biological Sciences, Ingelheim (Canada) Ltd, Laval, QC, H7S 2G5, Can.
SOURCE: Nature (London, United Kingdom) (2003), 426(6963), 186-189
CODEN: NATUAS; ISSN: 0028-0836
PUBLISHER: Nature Publishing Group

DOCUMENT TYPE: Journal
LANGUAGE: English
REFERENCE COUNT: 30 THERE ARE 30 CITED REFERENCES
AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L14 ANSWER 3 OF 3 HCAPLUS COPYRIGHT 2004 ACS on STN
ACCESSION NUMBER: 2000:725652 HCAPLUS Full-text
DOCUMENT NUMBER: 133:296659
TITLE: Preparation of macrocyclic peptides active against the hepatitis C virus
INVENTOR(S): Tsantrizos, Youla S.; Cameron, Dale R.; Faucher, Anne-marie; Ghio, Elise; Goudreau, Nathalie; Halmos, Teddy; Llinas-brunet, Montse
PATENT ASSIGNEE(S): Boehringer Ingelheim (Canada) Ltd., Can.
SOURCE: PCT Int. Appl., 154 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO.
DATE -----

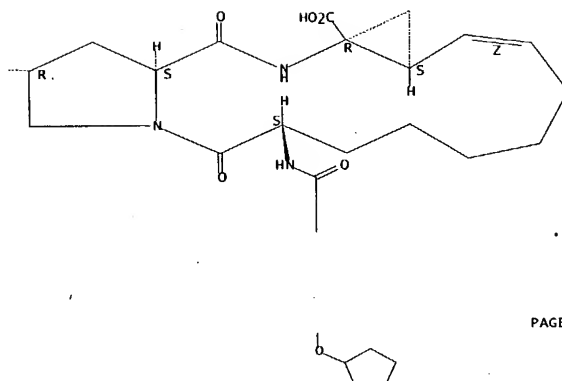
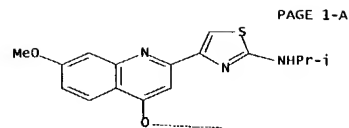
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 OTHER SOURCE(S): MARPAT 133:296659
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L14 ANSWER 1 OF 3 HCAPLUS COPYRIGHT 2004 ACS ON STN
 IT 300832-84-2, BILN 2061
 RL: THU (Therapeutic use); BTOL (Biological study); USES (Uses)
 (spiro compds. for inhibiting the first-pass effect)
 RN 300832-84-2 HCAPLUS
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 carboxylic
 acid, 6-[[[(cyclopentyl)oxy]carbonyl]amino]-
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Absolute stereochemistry.
 Double bond geometry as shown.



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 ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
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 F6 11 PCTFULL
 F7 10 CAPLUS
 F8 10 DDFU
 F9 10 DRUGU
 F10 8 BIOTECHNO
 F11 6 USPATFULL
 F12 4 LIFESCI
 F13 4 PASCAL
 F14 4 TOXCENTER
 F15 3 ESRIOBASE
 F16 3 IMSDRUGNEWS
 F17 3 NLDB
 F18 2 ADISCTI
 F19 2 BIOENG
 F20 2 CBNB
 F21 2 COMPENDEX
 F22 2 PHIN
 F23 2 WPIDS
 F24 2 WPINDEX
 F25 1 BABS
 F26 1 CIN
 F27 1 EMBAL
 F28 1 IFIPAT
 F29 1 PROMT

```
=> index f1-f29
COST IN U.S. DOLLARS          SINCE FILE
TOTAL                          ENTRY
SESSION
FULL ESTIMATED COST          1.14
94.22

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE
TOTAL                          ENTRY
SESSION
CA SUBSCRIBER PRICE          0.00
4.90

INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
CAPLUS, DDFU,
      DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
ESBIOBASE,
      IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
WPIDS,
      WPINDEX, BABS, CIN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20
ON 26 OCT 2004
```

29 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view
search error messages that display as 0* with SET DETAIL OFF.

=> d his 114-

```
(FILE 'HCAPLUS' ENTERED AT 14:32:46 ON 26 OCT 2004)
L14 3 L10 AND PD<20030327

FILE 'HOME' ENTERED AT 14:39:45 ON 26 OCT 2004

INDEX 'IMOBILITY, ZMOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA,
ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS,
BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS,
BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...' ENTERED AT
15:24:55 ON
26 OCT 2004
```

SEA BILN 2061

```
2 FILE ADISCTI
1 FILE BABS
2 FILE BIOENG
18 FILE BIOSIS
8 FILE BIOTECHNO
10 FILE CAPLUS
2 FILE CBNB
1 FILE CIN
```

```
20 "BILN"
1 "BILNS"
21 "BILN"
1864 "2061"
15 BILN 2061
("BILN"(w)"2061")
34452 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'MEDLINE'
19 BILN
122 2061
13 BILN 2061
(BILN(w)2061)
122732 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'PCTFULL'
40 BILN
2832 2061
11 BILN 2061
(BILN(w)2061)
127299 CRYSTAL?
9 L15 AND CRYSTAL?
FILE 'CAPLUS'
20 BILN
382 2061
10 BILN 2061
(BILN(w)2061)
1605797 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'DDFU'
13 BILN
21 2061
10 BILN 2061
(BILN(w)2061)
7961 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'DRUGU'
13 BILN
36 2061
10 BILN 2061
(BILN(w)2061)
12117 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'BIOTECHNO'
12 BILN
43 2061
8 BILN 2061
(BILN(w)2061)
27086 CRYSTAL?
3 L15 AND CRYSTAL?
FILE 'USPATFULL'
17 BILN
4661 2061
6 BILN 2061
(BILN(w)2061)
570101 CRYSTAL?
```

```
2 FILE COMPENDEX
10 FILE DDFU
10 FILE DRUGU
1 FILE EMBAL
42 FILE EMBASE
3 FILE ESBIOBASE
1 FILE IFIPAT
3 FILE IMSDRUGNEWS
15 FILE INVESTEXT
4 FILE LIFESCI
13 FILE MEDLINE
3 FILE NLDB
4 FILE PASCAL
11 FILE PCTFULL
2 FILE PHIN
1 FILE PROMT
22 FILE SCISEARCH
4 FILE TOXCENTER
6 FILE USPATFULL
2 FILE WPIDS
2 FILE WPINDEX
L15 QUE BILN 2061
-----
```

```
INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
CAPLUS,
DDFU, DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
ESBIOBASE,
IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
WPIDS,
WPINDEX, BABS, CIN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20 ON
26 OCT
2004
```

=> l15 and crystal?

```
FILE 'EMBASE'
48 "BILN"
124 "2061"
42 BILN 2061
("BILN"(w)"2061")
97097 CRYSTAL?
4 L15 AND CRYSTAL?
FILE 'SCISEARCH'
27 BILN
168 2061
22 BILN 2061
(BILN(w)2061)
689047 CRYSTAL?
3 L15 AND CRYSTAL?
FILE 'BIOSIS'
27 BILN
129 2061
18 BILN 2061
(BILN(w)2061)
111782 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'INVESTEXT'
```

```
6 L15 AND CRYSTAL?
FILE 'LIFESCI'
5 "BILN"
28 "2061"
4 BILN 2061
("BILN"(w)"2061")
27296 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'PASCAL'
13 BILN
69 2061
4 BILN 2061
(BILN(w)2061)
517682 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'TOXCENTER'
6 BILN
63 2061
4 BILN 2061
(BILN(w)2061)
66181 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'ESBIOBASE'
3 BILN
49 2061
3 BILN 2061
(BILN(w)2061)
41699 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'IMSDRUGNEWS'
3 "BILN"
5 "2061"
3 BILN 2061
("BILN"(w)"2061")
118 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'NLDB'
4 "BILN"
274 "2061"
3 BILN 2061
("BILN"(w)"2061")
32830 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'ADISCTI'
3 BILN
21 2061
2 BILN 2061
(BILN(w)2061)
463 CRYSTAL?
0 L15 AND CRYSTAL?
FILE 'BIOENG'
2 BILN
7 2061
2 BILN 2061
(BILN(w)2061)
7912 CRYSTAL?
0 L15 AND CRYSTAL?
```

```

FILE 'CBNB'
  2 BILN
  15 2061
  2 BILN 2061
    (BILN(W)2061)
  5823 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'COMPENDEX'
  4 BILN
  22 2061
  2 BILN 2061
    (BILN(W)2061)
  443469 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'PHIN'
  3 "BILN"
  35 "2061"
  2 BILN 2061
    ("BILN"(W)"2061")
  1251 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'WPIDS'
  4 BILN
  70 2061
  2 BILN 2061
    (BILN(W)2061)
  362675 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'WPINDEX'
  4 BILN
  70 2061
  2 BILN 2061
    (BILN(W)2061)
  362675 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'BABS'
  1 BILN
  24 2061
  1 BILN 2061
    (BILN(W)2061)
  91052 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'CIN'
  1 "BILN"
  12 "2061"
  1 BILN 2061
    ("BILN"(W)"2061")
  6321 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'EMBAL'
  1 BILN
  4 2061
  1 BILN 2061
    (BILN(W)2061)
  1192 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'IFIPAT'

```

```

=> L16
L18      25 L16

=> dup rem
ENTER L# LIST OR (END):L18
PROCESSING COMPLETED FOR L18
L19      21 DUP REM L18 (4 DUPLICATES REMOVED)

```

```

=> L19 and pd<20030327
  1 FILES SEARCHED...
  3 FILES SEARCHED...
L20      2 L19 AND PD<20030327

```

```

=> d 120 1-2 ibib hitstr abs kwic
'HITSTR' IS NOT A VALID FORMAT FOR FILE 'EMBASE'

```

The following are valid formats:

The default display format is BIB.

```

ABS ----- AB
ALL ----- AN, DN, TI, AU, CS, SO, PUI, CY, DT, FS, LA, SL, AB,
           CT, RN, CN, NP, CO, GEN
BIB ----- AN, DN, TI, AU, CS, SO, PUI, CY, DT, FS, LA, SL
CBIB ----- Compressed bibliographic data
DALL ----- ALL, delimited for post-processing
IABS ----- ABS, with a text label
IALL ----- ALL, indented with text labels
IBIB ----- BIB, indented with text labels
IND ----- CT, RN, CN, NP, CO, GEN
TRIAL ----- TI, CT, RN, CN, NP, CO, GEN
              (SAM, TRI)
HIT ----- All fields containing hit terms
HITIND ----- IND
KWIC ----- All hit terms plus 20 words on either side
OCC ----- List of display fields containing hit terms
              and number of occurrences in each field

```

Hit terms will be highlighted in all displayable fields.

To display a particular field or fields, enter the display field codes. For a list of display field codes, enter 'HELP DFIELDS' at an arrow prompt (=>). Examples of formats include: 'BIB'; 'AB'; 'SO ST'. You may specify the format fields in any order, and the information will be displayed in the same order as the format specification.

The same formats (except for HIT, HITIND, KWIC, and OCC) may be used with the DISPLAY ACC command to display the record for a specified Accession Number.

ENTER DISPLAY FORMAT (BIB):all

```

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```

```

  1 BILN
  39 2061
  1 BILN 2061
    (BILN(W)2061)
  146324 CRYSTAL?
    0 L15 AND CRYSTAL?
FILE 'PROMT'
  7 "BILN"
  538 "2061"
  1 BILN 2061
    ("BILN"(W)"2061")
  95992 CRYSTAL?
    0 L15 AND CRYSTAL?

```

L16 QUE L15 AND CRYSTAL?

```

=> d rank
F1      9 PCTFULL
F2      6 USPATFULL
F3      4 EMBASE
F4      3 SCISEARCH
F5      3 BIOTECHNO

```

```

=> file f3-5,f1,f2
COST IN U.S. DOLLARS      SINCE FILE
TOTAL                      ENTRY

```

```

SESSION
FULL ESTIMATED COST      1.71
95.93

```

```

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE
TOTAL                      ENTRY

```

```

SESSION
CA SUBSCRIBER PRICE      0.00
4.90

```

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FILE 'PCTFULL' ENTERED AT 15:27:48 ON 26 OCT 2004
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FILE 'USPATFULL' ENTERED AT 15:27:48 ON 26 OCT 2004
CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

```

=> L16, dup rem
L17      0 L16, DUP REM

```

ACCESSION NUMBER: 2003468113 EMBASE Full-text
TITLE: Current therapy and new molecular approaches to
antiviral

AUTHOR: Hugle T.; Cerny A.
CORPORATE SOURCE: Dr. A. Cerny, Clinica Medica, Ospedale Civico,
CH-6903

SOURCE: Lugano, Switzerland. andreas.cerny@bluewin.ch
Reviews in Medical Virology, (2003) 13/6
(361-371).

Refs: 79
ISSN: 1052-9276 CODEN: RMVIEW

COUNTRY: United Kingdom
DOCUMENT TYPE: Journal; General Review
FILE SEGMENT: 004 Microbiology
030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
039 Pharmacy

LANGUAGE: English
SUMMARY LANGUAGE: English

ABSTRACT:
Current therapeutic options for hepatitis C are limited, especially for genotype 1. For genotypes 2 and 3, pegylated interferon in combination with ribavirin, can lead to a sustained virological response in up to 80% of patients. Unfortunately, adverse effects of IFN and ribavirin are a major problem and the list of contraindications for HCV therapy is long, including decompensated cirrhosis of the liver and psychiatric disorders. Therefore, alternative therapeutic approaches are needed. New delivery options for IFN and ribavirin are aimed at optimising efficiency and reducing adverse effects. Recent progress in the molecular virology of HCV has identified new targets for antiviral intervention. Inhibition of HCV gene expression and replication as well as immunotherapeutic concepts aimed at enhancing the cellular response against HCV are being explored. Solution of the crystal structures of HCV key enzymes led to the design of specific inhibitors including compounds active against the well characterised NS3 serine protease and RNA-dependent RNA polymerase which are currently in the early phase clinical investigation. New strategies for inhibiting HCV gene expression include the use of antisense oligodeoxynucleotides and ribozymes. Immunomodulation by agents such as inosine monophosphate dehydrogenase inhibitors, thymosin- α 1, histamine or amantadine are being

studied in combination with IFN and/or ribavirin. Immunotherapeutic vaccination with recombinant HCV E1 protein improved host immunity against HCV and thus seems to be a promising new option. Copyright .COPYRG. 2003 John Wiley & Sons, Ltd.

CONTROLLED TERM: Medical Descriptors:
 *hepatitis C: DT, drug therapy
 *hepatitis C: ET, etiology
 *hepatitis C: PC, prevention
 *infection prevention
 virus gene
 genotype
 drug response
 drug contraindication
 drug delivery system
 side effect: SI, side effect
 gene expression
 drug targeting
 immunotherapy
 enzyme structure
 crystal structure
 drug design
 drug activity
 antiviral activity
 protein targeting
 immunomodulation
 vaccination
 Hepatitis C virus
 immune response
 cellular immunity
 hemolytic anemia: SI, side effect
 mental disease: SI, side effect
 flu like syndrome: SI, side effect
 leukopenia: SI, side effect
 thrombocytopenia: SI, side effect
 teratogenicity
 virus replication
 drug hypersensitivity: SI, side effect
 rash: SI, side effect
 human
 nonhuman
 clinical trial
 review
 Drug Descriptors:
 alpha interferon: AE, adverse drug reaction
 alpha interferon: CT, clinical trial
 alpha interferon: CB, drug combination
 alpha interferon: DT, drug therapy
 alpha interferon: TO, drug toxicity
 alpha interferon: PR, pharmaceuticals
 alpha interferon: PD, pharmacology
 alpha interferon: SC, subcutaneous drug
 administration
 ribavirin: AE, adverse drug reaction

trial

combination

therapy

pharmacology

CAS REGISTRY NO.: 37205-61-1;

198821-22-6,

CHEMICAL NAME:

COMPANY NAME:

NABI;

ribozyme: CT, clinical trial
 ribozyme: DT, drug therapy
 ribozyme: TO, drug toxicity
 ribozyme: PD, pharmacology
 hepatocyte: AE, adverse drug reaction
 hepatocyte: CT, clinical trial
 hepatocyte: DT, drug therapy
 hepatocyte: TO, drug toxicity
 hepatocyte: PD, pharmacology
 antisense oligodeoxynucleotide: CT, clinical
 antisense oligodeoxynucleotide: DT, drug therapy
 antisense oligodeoxynucleotide: PD, pharmacology
 isis 14803: CT, clinical trial
 isis 14803: DT, drug therapy
 isis 14803: PD, pharmacology
 RNA derivative: DV, drug development
 RNA derivative: DT, drug therapy
 RNA derivative: PD, pharmacology
 small interfering rna: DV, drug development
 small interfering rna: DT, drug therapy
 small interfering rna: PD, pharmacology
 monoclonal antibody: DT, drug therapy
 monoclonal antibody: PD, pharmacology
 xtl 002: DT, drug therapy
 xtl 002: PD, pharmacology
 cicavir: DT, drug therapy
 cicavir: PD, pharmacology
 immunomodulating agent: CB, drug combination
 immunomodulating agent: DT, drug therapy
 thymosin alpha1: CT, clinical trial
 thymosin alpha1: CB, drug combination
 thymosin alpha1: DO, drug dose
 thymosin alpha1: DT, drug therapy
 thymosin alpha1: PD, pharmacology
 inosinate dehydrogenase inhibitor: CB, drug
 inosinate dehydrogenase inhibitor: DT, drug
 inosinate dehydrogenase inhibitor: PD,
 merimepodib: CT, clinical trial
 merimepodib: CB, drug combination
 merimepodib: DT, drug therapy
 merimepodib: PD, pharmacology
 unindexed drug
 unclassified drug
 (ribavirin) 36791-04-5; (proteinase inhibitor)
 (thymosin alpha1) 69521-94-4; (merimepodib)
 198821-38-4
 (1) vx 950; (2) jtk 003; Biln 2061; Isis 14803;
 xtl 002
 (1) Vertex; (2) Akros; Ribozyme Pharmaceuticals;
 Sciclone; Regenerx; Maxim

ribavirin: CT, clinical trial
 ribavirin: CB, drug combination
 ribavirin: CM, drug comparison
 ribavirin: DT, drug therapy
 ribavirin: PK, pharmacokinetics
 ribavirin: PD, pharmacology
 ribavirin: PO, oral drug administration
 albumin conjugate: PR, pharmaceuticals
 liposome: PR, pharmaceuticals
 polyaminoacid: PR, pharmaceuticals
 polyaminoacid: PO, oral drug administration
 ribavirin derivative: AE, adverse drug reaction
 ribavirin derivative: CT, clinical trial
 ribavirin derivative: CB, drug combination
 ribavirin derivative: CM, drug comparison
 ribavirin derivative: DT, drug therapy
 ribavirin derivative: PD, pharmacology
 viramidine: AE, adverse drug reaction
 viramidine: CT, clinical trial
 viramidine: CB, drug combination
 viramidine: CM, drug comparison
 viramidine: DT, drug therapy
 viramidine: PD, pharmacology
 levovirin: AE, adverse drug reaction
 levovirin: CT, clinical trial
 levovirin: CM, drug comparison
 levovirin: DT, drug therapy
 levovirin: PD, pharmacology
 proteinase inhibitor: AE, adverse drug reaction
 proteinase inhibitor: CT, clinical trial
 proteinase inhibitor: DO, drug dose
 proteinase inhibitor: DT, drug therapy
 proteinase inhibitor: PK, pharmacokinetics
 proteinase inhibitor: PD, pharmacology
 proteinase inhibitor: PO, oral drug
 administration
 biln 2061: AE, adverse drug reaction
 biln 2061: CT, clinical trial
 biln 2061: DO, drug dose
 biln 2061: DT, drug therapy
 biln 2061: PK, pharmacokinetics
 biln 2061: PD, pharmacology
 biln 2061: PO, oral drug administration
 vx 950: DT, drug therapy
 vx 950: PD, pharmacology
 virus protein
 protein NS5B
 RNA directed DNA polymerase inhibitor: CT,
 clinical trial
 RNA directed DNA polymerase inhibitor: DT, drug
 therapy
 RNA directed DNA polymerase inhibitor: PD,
 pharmacology
 jtk 003: CT, clinical trial
 jtk 003: DT, drug therapy
 jtk 003: PD, pharmacology
 ribozyme: AE, adverse drug reaction

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 on STN
 ACCESSION NUMBER: 2003195244 EMBASE Full-text
 TITLE: Hepatitis C virus therapies: Current treatments, targets and future perspectives.
 AUTHOR: Walker M.P.; Appleby T.C.; Zhong W.; Lau J.Y.N.; Hong Z.
 CORPORATE SOURCE: Z. Hong, Ribapharm Inc., Hyland Avenue, Costa Mesa, CA, United States. zhihong@ribapharm.com
 SOURCE: Antiviral Chemistry and Chemotherapy, (2003) 14/1 (1-21).
 Refs: 208
 ISSN: 0956-3202 CODEN: ACCHEH
 COUNTRY: United Kingdom
 DOCUMENT TYPE: Journal; General Review
 FILE SEGMENT: 004 Microbiology
 030 Pharmacology
 037 Drug Literature Index
 038 Adverse Reactions Titles
 048 Gastroenterology
 LANGUAGE: English
 SUMMARY LANGUAGE: English
 ABSTRACT: Chronic hepatitis C virus (HCV) infection is the cause of an emerging global epidemic of chronic liver disease. Current combination therapies are at best 80% efficacious and are often poorly tolerated. Strategies to improve the therapeutic response include the development of novel interferons, nucleoside analogues with reduced haemolysis compared with ribavirin and inosine 5'-monophosphate dehydrogenase inhibitors. Compounds in preclinical or early clinical trials include small molecules that inhibit virus-specific enzymes (such as the serine proteases, RNA polymerase and helicase) or interfere with translation (including antisense molecules, iRNA and ribozymes). Advances in understanding HCV replication, obtaining a sub-genomic replicon and contriving potential small animal models, in addition to solving crystallographic structures for the replication enzymes, have improved prospects for developing novel therapies. This review summarizes current and evolving treatments for chronic hepatitis C infection. In addition, progress in HCV targets and drug discovery tools valuable in the search for novel anti-HCV agents is detailed.

CONTROLLED TERM:

Medical Descriptors:
 *hepatitis C: DT, drug therapy
 *hepatitis C: EP, epidemiology
 *hepatitis C: ET, etiology
 *chronic liver disease: ET, etiology
 drug efficacy
 drug tolerance
 hemolytic anemia: SI, side effect
 side effect: SI, side effect
 alanine aminotransferase blood level
 virus replication
 replicon
 crystal structure
 RNA translation
 untranslated region
 internal ribosome entry site
 monotherapy
 virus load
 treatment outcome
 treatment indication
 immunomodulation
 drug safety
 treatment failure
 chimpanzee
 transgenic mouse
 Hepatitis GB virus B
 IC 50
 structure activity relation
 drug structure
 virus assembly
 human
 nonhuman
 clinical trial
 review
 priority journal
 Drug Descriptors:
 *antivirus agent: AE, adverse drug reaction
 *antivirus agent: CT, clinical trial
 *antivirus agent: AN, drug analysis
 *antivirus agent: CB, drug combination
 *antivirus agent: CM, drug comparison
 *antivirus agent: DV, drug development
 *antivirus agent: DO, drug dose
 *antivirus agent: DT, drug therapy
 *antivirus agent: PD, pharmacology
 *antivirus agent: IV, intravenous drug
 *antivirus agent: SC, subcutaneous drug
 alpha interferon: AE, adverse drug reaction
 alpha interferon: CB, drug combination
 alpha interferon: CM, drug comparison
 alpha interferon: DO, drug dose
 alpha interferon: DT, drug therapy
 alpha interferon: PD, pharmacology
 nucleoside derivative: AN, drug analysis
 nucleoside derivative: CM, drug comparison

administration

administration

levovirin: DV, drug development
 levovirin: DO, drug dose
 levovirin: DT, drug therapy
 levovirin: PD, pharmacology
 viramidine: CT, clinical trial
 viramidine: AN, drug analysis
 viramidine: CM, drug comparison
 viramidine: DV, drug development
 viramidine: DO, drug dose
 viramidine: DT, drug therapy
 viramidine: PD, pharmacology
 merimepodib: CT, clinical trial
 merimepodib: AN, drug analysis
 merimepodib: CB, drug combination
 merimepodib: CM, drug comparison
 merimepodib: DV, drug development
 merimepodib: DT, drug therapy
 merimepodib: PD, pharmacology
 thymosin alpha1: CT, clinical trial
 thymosin alpha1: AN, drug analysis
 thymosin alpha1: CB, drug combination
 thymosin alpha1: DV, drug development
 thymosin alpha1: DO, drug dose
 thymosin alpha1: DT, drug therapy
 thymosin alpha1: PD, pharmacology
 thymosin alpha1: SC, subcutaneous drug

administration

amantadine: CT, clinical trial
 amantadine: AN, drug analysis
 amantadine: CB, drug combination
 amantadine: CM, drug comparison
 amantadine: DV, drug development
 amantadine: PD, pharmacology
 recombinant interleukin 12: CT, clinical trial
 recombinant interleukin 12: AN, drug analysis
 recombinant interleukin 12: CB, drug combination
 recombinant interleukin 12: CM, drug comparison
 recombinant interleukin 12: DV, drug development
 recombinant interleukin 12: DO, drug dose
 recombinant interleukin 12: DT, drug therapy
 recombinant interleukin 12: PD, pharmacology
 histamine: CT, clinical trial
 histamine: AN, drug analysis
 histamine: CB, drug combination
 histamine: DV, drug development
 histamine: DT, drug therapy
 histamine: PD, pharmacology
 gamma interferon: CT, clinical trial
 gamma interferon: AN, drug analysis
 gamma interferon: CB, drug combination
 gamma interferon: DV, drug development
 gamma interferon: DT, drug therapy
 gamma interferon: PD, pharmacology
 proteinase inhibitor: CT, clinical trial
 proteinase inhibitor: DO, drug dose
 proteinase inhibitor: PD, pharmacology
 proteinase inhibitor: PO, oral drug

nucleoside derivative: DV, drug development
 nucleoside derivative: PR, pharmaceuticals
 nucleoside derivative: PD, pharmacology
 ribavirin: AE, adverse drug reaction
 ribavirin: CT, clinical trial
 ribavirin: CB, drug combination
 ribavirin: CM, drug comparison
 ribavirin: DO, drug dose
 ribavirin: DT, drug therapy
 ribavirin: PD, pharmacology
 inosinate dehydrogenase inhibitor: CM, drug
 inosinate dehydrogenase inhibitor: DT, drug
 inosinate dehydrogenase inhibitor: PD,
 serine proteinase: EC, endogenous compound
 RNA polymerase: EC, endogenous compound
 helicase: EC, endogenous compound
 ribozyme: EC, endogenous compound
 recombinant alpha2a interferon: CM, drug
 recombinant alpha2a interferon: DO, drug dose
 recombinant alpha2a interferon: DT, drug therapy
 recombinant alpha2a interferon: PD, pharmacology
 recombinant alpha2a interferon: SC, subcutaneous
 administration
 recombinant alpha2b interferon: CM, drug
 recombinant alpha2b interferon: DO, drug dose
 recombinant alpha2b interferon: DT, drug therapy
 recombinant alpha2b interferon: PD, pharmacology
 recombinant alpha2b interferon: SC, subcutaneous
 administration
 consensus interferon: CM, drug comparison
 consensus interferon: DO, drug dose
 consensus interferon: DT, drug therapy
 consensus interferon: PD, pharmacology
 consensus interferon: SC, subcutaneous drug
 peginterferon alpha2b: CT, clinical trial
 peginterferon alpha2b: CB, drug combination
 peginterferon alpha2b: CM, drug comparison
 peginterferon alpha2b: DO, drug dose
 peginterferon alpha2b: DT, drug therapy
 peginterferon alpha2b: PD, pharmacology
 peginterferon alpha2a: CT, clinical trial
 peginterferon alpha2a: CB, drug combination
 peginterferon alpha2a: CM, drug comparison
 peginterferon alpha2a: DO, drug dose
 peginterferon alpha2a: DT, drug therapy
 peginterferon alpha2a: PD, pharmacology
 levovirin: CT, clinical trial
 levovirin: AN, drug analysis
 levovirin: CM, drug comparison

comparison

therapy

pharmacology

comparison

drug

comparison

drug

administration

administration

biln 2061: CT, clinical trial
 biln 2061: DO, drug dose
 biln 2061: PD, pharmacology
 biln 2061: PO, oral drug administration
 peptide derivative: AN, drug analysis
 peptide derivative: DV, drug development
 peptide derivative: PD, pharmacology
 peptide alpha keto acid: AN, drug analysis
 peptide alpha keto acid: DV, drug development
 peptide alpha keto acid: PD, pharmacology
 pyrrolidine derivative: AN, drug analysis
 pyrrolidine derivative: DV, drug development
 pyrrolidine derivative: PD, pharmacology
 pyrrolidine 5,5 lactam: AN, drug analysis
 pyrrolidine 5,5 lactam: DV, drug development
 pyrrolidine 5,5 lactam: PD, pharmacology
 IDb3: DV, drug development
 IDb3: PD, pharmacology
 unindexed drug
 unclassified drug
 isis 14803
 gw 3112
 gw 2549
 gw 0569
 n [4 [[[[6,7 dihydro 2 (4 methylphenyl) 5h
 8 yl[carbonyl]amino]benzyl] n,n dimethyl 2h
 4 aminium chloride
 1,1' [1,4 phenylenebis(methylene)]bis(1,4,8,11
 tetraazacyclotetradecane)
 (ribavirin) 36791-04-5; (serine proteinase)
 (RNA polymerase) 9014-24-8; (helicase) 42613-29-
 6;
 (recombinant alpha2b interferon) 98530-12-2;
 alpha2b) 215647-85-1; (peginterferon alpha2a)
 (merimepodib) 198821-22-6, 198821-38-4; (thymosin
 alpha1)
 69521-94-4; (amantadine) 665-66-7, 768-94-5;
 (histamine)
 51-45-6, 56-92-8, 93443-21-1; (gamma interferon)
 82115-62-6; (proteinase inhibitor) 37205-61-1; (n
 [4 [[[[6,7
 dihydro 2 (4 methylphenyl) 5h benzocycloheptan 8
 yl[carbonyl]amino]benzyl] n,n dimethyl 2h
 tetrahydropyran 4
 aminium chloride) 229005-80-5; (1,1' [1,4
 phenylenebis(methylene)]bis(1,4,8,11
 tetraazacyclotetradecane)) 155148-31-5
 (1) vx 497; (2) ceplene; (3) biln 2061; (4) isis
 14803; Zadaxin; GW 3112; GW 2549; GW 0569; Tak
 3100; IDb3

benzocycloheptan

tetrahydropyran

CAS REGISTRY NO.:

6;

(peginterferon

198153-51-4;

alpha1)

(histamine)

[4

tetrahydropyran 4

CHEMICAL NAME:

779; Amd

COMPANY NAME: (1) Vertex (United States); (2) Maxim; (3)
Boehringer Ingetheim; (4) Isis (United States); Ribapharm;
Merck (United States); Glaxo SmithKline (United
Kingdom); Bristol Myers Squibb (United States); Celera (United
States); Viropharma; Japanese tobacco; IRBM

=> d his

(FILE 'HOME' ENTERED AT 14:24:22 ON 26 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE,
AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS,
BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN,
CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 14:25:23 ON
26 OCT 2004

L1 FILE 'REGISTRY' ENTERED AT 14:26:32 ON 26 OCT 2004

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SEL RN
SET SMA LOGIN

INDEX 'BEILSTEIN, GELIN' ENTERED AT 14:27:43 ON 26 OCT 2004
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L2 QUE 300832-84-2/BI AND OPTICAL?/FA

L3 FILE 'CAPLUS' ENTERED AT 14:28:02 ON 26 OCT 2004
0 S L1/BMF

L4 FILE 'CAPLUS' ENTERED AT 14:28:18 ON 26 OCT 2004
0 S L1/BPN

L5 FILE 'CAPLUS' ENTERED AT 14:28:28 ON 26 OCT 2004
0 S L1/IMF

L6 FILE 'CAPLUS' ENTERED AT 14:28:41 ON 26 OCT 2004
1 S L1/PEP

L7 FILE 'CAPLUS' ENTERED AT 14:29:30 ON 26 OCT 2004
0 S L1/PUR

L8 FILE 'CAPLUS' ENTERED AT 14:29:48 ON 26 OCT 2004
6 S (L1/SPN OR L1/CPN)
L9 6 FOCUS L8 1-

FILE 'REGISTRY' ENTERED AT 14:31:45 ON 26 OCT 2004

IMSDRUGNEWS, NLDB, ADISCTI, BIOENG, CBNB, COMPENDEX, PHIN,
WPIOS, WPINDEK, BABS, CTN, EMBAL, IFIPAT, PROMT' ENTERED AT 15:26:20 ON
26 OCT 2004

SEA L15 AND CRYSTAL?

4 FILE EMBASE
3 FILE SCISEARCH
9 FILE PCTFULL
3 FILE BIOTECHNO
6 FILE USPATFULL
L16 QUE L15 AND CRYSTAL?

AT FILE 'EMBASE, SCISEARCH, BIOTECHNO, PCTFULL, USPATFULL' ENTERED

15:27:48 ON 26 OCT 2004

L17 0 L16, DUP REM

L18 25 L16

L19 21 DUP REM L18 (4 DUPLICATES REMOVED)

L20 2 L19 AND PD<20030327

=> stindex

ENTER FILE OR CLUSTER NAMES (NONE):a11
FILE 'ENCOMPLIT' ACCESS NOT AUTHORIZED
FILE 'ENCOMPLIT2' ACCESS NOT AUTHORIZED
FILE 'ENCOMPPAT' ACCESS NOT AUTHORIZED
FILE 'ENCOMPPAT2' ACCESS NOT AUTHORIZED
COST IN U.S. DOLLARS

TOTAL SINCE FILE

SESSION ENTRY

FULL ESTIMATED COST 16.52

112.45

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE

TOTAL ENTRY

SESSION 0.00

CA SUBSCRIBER PRICE

4.90

INDEX 'IMOBILITY, 2MOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA, ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS, BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS, BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...'
ENTERED AT 15:32:46 ON 26 OCT 2004

143 FILES IN THE FILE LIST IN STINDEX

Enter SET DETAIL ON to see search term postings or to view
search error messages that display as 0* with SET DETAIL OFF.

FILE 'HCAPLUS' ENTERED AT 14:32:46 ON 26 OCT 2004

L10 15 S L1
L11 0 L10 AND CRYSTAL
L12 0 L10 AND CRYSTAL?
L13 0 L10 AND ALCOHOL
L14 3 L10 AND PD<20030327

FILE 'HOME' ENTERED AT 14:39:45 ON 26 OCT 2004

INDEX 'IMOBILITY, 2MOBILITY, ABI-INFORM, ADISCTI, AEROSPACE,
AGRICOLA, ALUMINIUM, ANABSTR, ANTE, APOLLIT, AQUALINE, AQUASCI, AQUIRE,
BABS, BIBLIODATA, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS,
BIOTECHABS, BIOTECHDS, BIOTECHNO, BLDB, CABA, CANCERLIT, ...' ENTERED AT
15:24:55 ON
26 OCT 2004

SEA BILN 2061

2 FILE ADISCTI
1 FILE BABS
2 FILE BIOENG
18 FILE BIOSIS
8 FILE BIOTECHNO
10 FILE CAPLUS
2 FILE CBNB
1 FILE CIN
2 FILE COMPENDEX
10 FILE DDFU
10 FILE DRUGU
1 FILE EMBAL
42 FILE EMBASE
3 FILE ESBIOBASE
1 FILE IFIPAT
3 FILE IMSDRUGNEWS
15 FILE INVESTEXT
4 FILE LIFESCI
13 FILE MEDLINE
3 FILE NLDB
4 FILE PASCAL
11 FILE PCTFULL
2 FILE PHIN
1 FILE PROMT
22 FILE SCISEARCH
4 FILE TOXCENTER
6 FILE USPATFULL
2 FILE WPIOS
2 FILE WPIINDEX
L15 QUE BILN 2061

INDEX 'EMBASE, SCISEARCH, BIOSIS, INVESTEXT, MEDLINE, PCTFULL,
CAPLUS, DDFU, DRUGU, BIOTECHNO, USPATFULL, LIFESCI, PASCAL, TOXCENTER,
ESBIOBASE,

=> ciluprevir/cn

FILE 'IMOBILITY'
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0 CILUPREVIR/CN

FILE '2MOBILITY'
'CN' IS NOT A VALID FIELD CODE
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FILE 'ABI-INFORM'
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FILE 'ADISCTI'
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FILE 'AGRICOLA'
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FILE 'ANABSTR'
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FILE 'BIOCOMMERCE'
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-----User Break-----
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FILE 'BIOSIS'
'CN' IS NOT A VALID FIELD CODE
0 CILUPREVIR/CN

SEARCH ENDED BY USER
FILE 'BIOTECHABS'
SEARCH ENDED BY USER


```
=> ciluprevir/
FILE 'IMOBILITY'
'CILUPREVIR/' IS NOT A VALID FIELD CODE
For a list of field codes for the current file, enter "HELP SFIELDS"
at an arrow prompt (=>).
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=> ciluprevir
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FILE '2MOBILITY'
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L21 QUE CILUPREVIR

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F1	3	EMBASE
F2	2	ESBIOBASE
F3	2	MEDLINE
F4	2	SCISEARCH
F5	1	DDFU
F6	1	DRUGU

=> fil f1-6		
COST IN U.S. DOLLARS		SINCE FILE
TOTAL		ENTRY
SESSION		
FULL ESTIMATED COST	5.70	
118.15		
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)		SINCE FILE
TOTAL		ENTRY
SESSION		
CA SUBSCRIBER PRICE	0.00	
4.90		
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FILE 'DDFU' ACCESS NOT AUTHORIZED

FILE 'DRUGU' ENTERED AT 15:39:01 ON 26 OCT 2004
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=> s 121
L22 10 L21

=> 122 and pd<20030327
1 FILES SEARCHED...
2 FILES SEARCHED...
'20030327' NOT A VALID FIELD CODE
'20030327' NOT A VALID FIELD CODE
L23 1 L22 AND PD<20030327

=> d 123

L23 ANSWER 1 OF 1 EMBASE COPYRIGHT 2004 ELSEVIER INC. ALL RIGHTS RESERVED.

on STN
AN 2004039472 EMBASE Full-text
TI Gateways to Clinical Trials.
AU Bayes M.; Rabasseda X.; Prous J.R.
CS M. Bayes, Prous Science, S.A., P.O. Box 540, 08080 Barcelona, Spain.
mbayes@prous.com
SO Methods and Findings in Experimental and Clinical Pharmacology,
(2003) 25/10 (831-855).
Refs: 145
ISSN: 0379-0355 CODEN: MFEPOX
CY Spain
DT Journal; General Review
FS 030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
LA English
SL English

=> FIL STNGUIDE
COST IN U.S. DOLLARS
TOTAL
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FULL ESTIMATED COST
126.52
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)
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CA SUBSCRIBER PRICE

SINCE FILE
ENTRY
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exenatide, exisulind, ezetimib; Febuxostat; Gallium maltolate, ganirelix acetate, garenoxacin mesilate, gefitinib; H11, HUMax; IL-15, IDD-1, IGIV-C, imatinib mesylate, ISIS-14803, ITF-1697, ivabradine hydrochloride; KRN-5500; L-365260, levetiracetam, levosimendan, licofelone, linezolid, LJP-1082, lopinavir, lumiracoxib; MCC-478, melatonin, morphine hydrochloride, morphine-6-glucuronide, moxidectin; N-Acetylcarnosine, natalizumab, NM-702, NNC-05-1869, NSC-703940; Ocinaclone OM-89, omalizumab, omeprazole/sodium bicarbonate, OPC-28326, ospemifene; PEG-filgrastim peginterferon alfa-2a, pegsunercept, pirfenidone, pralmorelin, pregabalin; Recombinant glucagon-like peptide-1 (7-36) amide, repifermin, RSD-1235; S-8184, selodexon, sodium dichloroacetate, suberanilohydroxamic acid; TAS-102, terfenadine, teriparatide, tipranavir troxatidine; ximelagatran; YM-337. .COPYRG. 2003 Prous Science. All rights reserved.

CT Medical Descriptors:
*drug monitoring
drug indication
drug efficacy
drug safety
side effect: SI, side effect
patient compliance
drug tolerability
liver toxicity: SI, side effect
bleeding: SI, side effect
disease exacerbation
systemic lupus erythematosus: SI, side effect
digestive system ulcer: SI, side effect
neutropenia: SI, side effect
teratogenicity: SI, side effect
human
clinical trial
review
Drug Descriptors:
abietimus: CT, clinical trial
abietimus: IV, intravenous drug administration
adalimumab: AE, adverse drug reaction
adalimumab: CT, clinical trial
linezolid: CT, clinical trial
alemtuzumab: CT, clinical trial
ivabradine: CT, clinical trial
ivabradine: IV, intravenous drug administration
recombinant interleukin 1 receptor blocking agent: CT, clinical trial
trial
recombinant interleukin 1 receptor blocking agent: IA,
intraarterial drug
administration
glucagon like peptide 1: CT, clinical trial
glucagon like peptide 1: SC, subcutaneous drug administration
astemizole: CT, clinical trial
atazanavir: CT, clinical trial
bosentan: CT, clinical trial
botulinum toxin B: CT, clinical trial
caspofungin: CT, clinical trial
ciclesonide: CT, clinical trial
cilomilast: CT, clinical trial

4.90

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on STN
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TI Gateways to Clinical Trials.
AU Bayes M.; Rabasseda X.; Prous J.R.
CS M. Bayes, Prous Science, S.A., P.O. Box 540, 08080 Barcelona, Spain.
mbayes@prous.com
SO Methods and Findings in Experimental and Clinical Pharmacology,
(2003) 25/10 (831-855).
Refs: 145
ISSN: 0379-0355 CODEN: MFEPOX
CY Spain
DT Journal; General Review
FS 030 Pharmacology
037 Drug Literature Index
038 Adverse Reactions Titles
LA English
SL English
AB Gateways to Clinical Trials is a guide to the most recent clinical trials in current literature and congresses. The data in the following tables has been retrieved from the Clinical Studies Knowledge Area of Prous Science Integrity*, the drug discovery and development portal, <http://integrity.prous.com>. This issue focuses on the following selection of drugs: Abietimus sodium, adalimumab, alefacept, alemtuzumab, almotriptan, AMGN-0007, anakinra, anti-CTLA-4 Mab, L-arginine hydrochloride, arzoifene hydrochloride, astemizole, atazanavir sulfate, atlizumab; Belimumab, BG-9928, binodenoson, bosentan, botulinum toxin type B, bovine lactoferrin, BufferGel; Caspofungin acetate, ciclesonide, cilomilast, ciluprevir, clofarabine, CVT-3146; Darbepoetin alfa, desloratadine, diflomotecan, doripenem, dronedarone hydrochloride, drotrecogin alfa (activated), DT388-GM-CSF, duloxetine hydrochloride, E-5564, efalizumab, enfuvirtide, esomeprazole magnesium, estradiol acetate, ETC-642,

efalizumab: CT, clinical trial
imatinib: CT, clinical trial
terfenadine: CT, clinical trial
tipranavir: CT, clinical trial
tipranavir: PO, oral drug administration
ximelagatran: CT, clinical trial
ximelagatran: PO, oral drug administration
ym 337: CT, clinical trial
moxidectin: CT, clinical trial
estradiol: CT, clinical trial
novel erythropoiesis stimulating protein: CT, clinical trial
novel erythropoiesis stimulating protein: IV, intravenous drug administration
novel erythropoiesis stimulating protein: SC, subcutaneous drug administration
desloratadine: CT, clinical trial
desloratadine: PO, oral drug administration
diflomotecan: CT, clinical trial
diflomotecan: IV, intravenous drug administration
diflomotecan: PO, oral drug administration
morphine: CT, clinical trial
etiracetam: CT, clinical trial
doripenem: CT, clinical trial
duloxetine: CT, clinical trial
unindexed drug
(abietimus) 167362-48-3, 169147-32-4; (adalimumab) 331731-18-1;
(linezolid)
165800-03-3; (alemtuzumab) 216503-57-0; (ivabradine) 148849-67-6,
148870-80-8, 155974-00-8; (glucagon like peptide 1) 89750-14-1;
(astemizole) 68844-77-9; (atazanavir) 198904-31-3; (bosentan)
147536-97-8,
157212-55-0; (caspofungin) 189768-38-5; (ciclesonide) 126544-47-6;
(cilomilast) 153259-65-5; (efalizumab) 214745-43-4; (imatinib)
152459-95-5, 220127-57-1; (terfenadine) 50679-08-8; (tipranavir)
174484-41-4; (ximelagatran) 192939-46-1, 260790-58-7;
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(etiracetam)
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116539-59-4, 136434-34-9
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L1 1 S 300832-84-2/RN
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SEA E1 AND OPTICAL7/FA

L2 QUE 300832-84-2/BI AND OPTICAL7/FA

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L9 6 FOCUS L8 1-

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L10 15 S L1
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L12 0 L10 AND CRYSTAL?
L13 0 L10 AND ALCOHOL
L14 3 L10 AND PD<20030327

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2 FILE ADISCTI

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4 FILE PASCAL
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6 FILE USPATFULL
2 FILE WPIDS
2 FILE WPINDEX
L15 QUE BILN 2061

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SEA L15 AND CRYSTAL?

4 FILE EMBASE
3 FILE SCISEARCH
9 FILE PCTFULL
3 FILE BIOTECHNO
6 FILE USPATFULL
L16 QUE L15 AND CRYSTAL?

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L17 0 L16, DUP REM
L18 25 L16
L19 21 DUP REM L18 (4 DUPLICATES REMOVED)
L20 2 L19 AND PD<20030327

=>
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COST IN U.S. DOLLARS	SINCE FILE
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FULL ESTIMATED COST	0.12
130.58	
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE
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New Data on BILN 2061, Experimental HCV Protease Inhibitor from ...

December 14-18, 2003. Kauai, Hawaii. New Data on **BILN 2061**, Experimental HCV Protease Inhibitor from Boehringer Ingelheim By Mark Nelson, MD. By Mark Nelson. ...

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BILN 2061 Establishes Proof-of-Concept in Humans for an HCV ...

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Yahoo! News - BILN 2061 Could Offer New Hope to Patients With ...

... Health - Acurian. **BILN 2061** Could Offer New Hope to Patients With Hepatitis C. Mon Oct 27, 7:00 PM ET. Source: Acurian Inc. by: Darrin Kiessling. ...

dynamics.org/~altenber/cryo/HCV/BILN2061.Yahoo.html - 43k - [Cached](#) - [Similar pages](#)

Hepatitis C Vets, Orally available Hepatitis C Virus (HCV) ...

... Orally available Hepatitis C Virus (HCV) Protease Inhibitor (**BILN 2061**, Boehringer Ingelheim Pharma) Demonstrates Potent Anti-viral Activity in Persons ...

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ISMC 2004: Session 2A: The discovery of BILN 2061, an NS3 protease ...

Session 2A: Keynote lecture. The discovery of **BILN 2061**, an NS3 protease inhibitor with antiviral effects in humans infected with hepatitis C virus. ...

www.ismc2004.dk/index.php/Session_2A_The_discovery_of_B/258/0/ - 11k - [Cached](#) - [Similar pages](#)

54th Annual Meeting of

... **BILN 2061** Establishes Proof-of-Concept in Humans for an HCV Protease Inhibitor ... Note:

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BILN 2061 Inhibits HCV Genotype 2 and 3 Proteases in Vitro, ... This study examined the ability of **BILN 2061** to inhibit NS3 protease from genotypes 2 and 3. ...

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Clinical Care Options for Hepatitis - BILN 2061 shows promise in ...

BILN 2061 shows promise in early clinical trial against HCV genotype 1,

Deanna M. Green, PhD. November 18, 2003 — **BILN 2061** induces ...

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Serin-Protease-Hemmstoff (BILN-2061) bei Hepatitis-C - [Translate this page]

... Medizinrecht. Neue Medikamente im Pipeline 2002. Serin-Protease-Hemmstoff (**BILN-2061**) bei Hepatitis-C. ... Die Arbeiten zu **BILN 2061** befinden sich am Anfangsphase. ...

www.medknowledge.de/neu/2002/IV-2002-32-biln-2061-pipeline.htm - 38k - [Cached](#) - [Similar pages](#)

HIV Report Jan 2003: A Promising New Anti-HCV Protease Inhibitor

... Researchers presented 4 papers describing the discovery, safety and early antiviral activity of **BILN 2061**, a serine protease inhibitor [Hepatology 2002;36:167A ...

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- ... COUNCIL: USAN **CILUPRE VIR**. PRONUNCIATION (si loo pre' veer). THERAPEUTIC CLAIM Treatment of Hepatitis C infection. CHEMICAL NAMES. 1 ...
- www.ama-assn.org/ama1/pub/upload/mm/365/ciluprevir.doc - Similar pages
- AMA (USAN) 2004 Published USAN
- ... atilmotin avanafil becatecarin bemotrizinol bisoctrizole canfosfamide hydrochloride cariporide mesylate ciclesonide **ciluprevir** dabuzalgron hydrochloride ...
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... However, the number of solutions required to describe the **crystallization protocol** is not limited to 2. Details of the **crystallization protocol** should be ...

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High Throughput Protein Crystallization

... with the aim to establish correlations between **crystallization** probability and ... Such an optimized **protocol** maximizes the probability of success while minimizing ...

[www-structure.llnl.gov/Xray/ tutorial/High_Throughput_EMBL_full.htm](http://www-structure.llnl.gov/Xray/tutorial/High_Throughput_EMBL_full.htm) - 79k - [Cached](#) - [Similar pages](#)

CRYSTOOL Move announcement

CRYSTOOL. Has Moved. There is a new implementation of Brent Segelke's CRYSTOOL program [1] for creating efficient, customizable, random **crystallization** screens. ...

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(IUCr) Crystallization experiments with 2-enoyl-CoA hydratase ...

... Volume 50 Part 4 Pages 443-447 July 1994 **Crystallization** experiments with 2-enoyl-CoA hydratase, using an automated 'fast-screening' **crystallization protocol**. ...

scripts.iucr.org/cgi-bin/paper?gr0322 - [Similar pages](#)

Protein Crystal Structure Team I | RIKEN

... (2), Development of high-throughput protein **crystallization protocol** for large scale.

(3), Development of in situ X-ray diffractometer for protein **crystal**. image ...

[www.riken.go.jp/engn/r-world/ research/lab/harima/group-h/crystal1/](http://www.riken.go.jp/engn/r-world/research/lab/harima/group-h/crystal1/) - 12k - [Cached](#) - [Similar pages](#)

Micro-seeding Procedure

... Pre-equilibrated drops contain protein and precipitant that is at a concentration just below what is required for precipitation or **crystallization** of the ...

www.hhmi.swmed.edu/Labs/rr/world/seeding.html - 3k - [Cached](#) - [Similar pages](#)

X-tal protocols: References.

... D50, 443-447 **Crystallization** Experiments with 2-Enoyl-CoA Hydratase, Using an Automated 'Fast-Screening' **Crystallization Protocol**. ...

www.xtal-protocols.de/ref/reference.html - 9k - [Cached](#) - [Similar pages](#)

Protein Crystallization

... Label your concentrated protein with a batch number. Differences in the purification **protocols** may later relate to irreproducibility in **crystallization**; ...

[www-structmed.cimr.cam.ac.uk/ Course/Crystals/Screening/hd_protocol.html](http://www-structmed.cimr.cam.ac.uk/Course/Crystals/Screening/hd_protocol.html) - 4k - [Cached](#) - [Similar pages](#)

The Scientist :: Crystal Illumination, Jan. 19, 2004

... usually 50 nanoliters, is so crucial to its drug-discovery efforts that the company has patented its proprietary Nano- volume **Crystallization™ protocol**. ...

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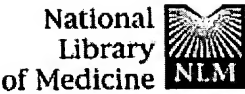
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#4	Search crystallization Field: All Fields , Limits: Review	09:47:07	1457
#3	Search crystallization	09:46:56	25455
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